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* * * * * * * * * * * * Welcome to STN International

NEWS 1 Web Page URLs for STN Seminar Sci

```
NEWS
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
                 "Ask CAS" for self-help around the clock
NEWS
         Jun 03
                 New e-mail delivery for search results now available
NEWS
         Aug 08
                 PHARMAMarketLetter(PHARMAML) - new on STN
         Aug 19
                 Aquatic Toxicity Information Retrieval (AQUIRE)
                 now available on STN
                 Sequence searching in REGISTRY enhanced
NEWS
         Aug 26
NEWS
         Sep 03
                 JAPIO has been reloaded and enhanced
NEWS
         Sep 16
                 Experimental properties added to the REGISTRY file
                 CA Section Thesaurus available in CAPLUS and CA
NEWS
         Sep 16
NEWS 10
                 CASREACT Enriched with Reactions from 1907 to 1985
         Oct 01
NEWS 11
         Oct 24
                 BEILSTEIN adds new search fields
NEWS 12
         Oct 24
                Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 13
         Nov 18
                 DKILIT has been renamed APOLLIT
         Nov 25 More calculated properties added to REGISTRY
NEWS 14
NEWS 15
        Dec 04
                 CSA files on STN
NEWS 16 Dec 17
                 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 17
         Dec 17
                 TOXCENTER enhanced with additional content
NEWS 18
         Dec 17
                 Adis Clinical Trials Insight now available on STN
NEWS 19
         Jan 29
                 Simultaneous left and right truncation added to COMPENDEX,
                 ENERGY, INSPEC
                 CANCERLIT is no longer being updated
NEWS 20
         Feb 13
NEWS 21
         Feb 24
                 METADEX enhancements
NEWS 22 Feb 24
                 PCTGEN now available on STN
NEWS 23 Feb 24
                 TEMA now available on STN
NEWS 24 Feb 26 NTIS now allows simultaneous left and right truncation
NEWS 25 Feb 26
                PCTFULL now contains images
NEWS 26 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 27 Mar 20
                EVENTLINE will be removed from STN
NEWS 28 Mar 24
                 PATDPAFULL now available on STN
NEWS 29 Mar 24
                Additional information for trade-named substances without
                 structures available in REGISTRY
NEWS 30
         Apr 11
                 Display formats in DGENE enhanced
NEWS 31
         Apr 14
                MEDLINE Reload
NEWS 32
         Apr 17
                 Polymer searching in REGISTRY enhanced
         Apr 21
NEWS 33
                 Indexing from 1947 to 1956 being added to records in CA/CAPLUS
NEWS 34
         Apr 21
                New current-awareness alert (SDI) frequency in
                 WPIDS/WPINDEX/WPIX
NEWS 35
        Apr 28
                RDISCLOSURE now available on STN
NEWS 36
                 Pharmacokinetic information and systematic chemical names
        May 05
                 added to PHAR
NEWS 37
                MEDLINE file segment of TOXCENTER reloaded
         May 15
                 Supporter information for ENCOMPPAT and ENCOMPLIT updated
NEWS 38
        May 15
                CHEMREACT will be removed from STN
NEWS 39
        May 16
        May 19 · Simultaneous left and right truncation added to WSCA
NEWS 40
        May 19 RAPRA enhanced with new search field, simultaneous left and
                 right truncation
```

MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS INTER General Internet Information

NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

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FILE 'HOME' ENTERED AT 13:36:13 ON 03 JUN 2003

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 13:36:28 ON 03 JUN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

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STRUCTURE FILE UPDATES: 1 JUN 2003 HIGHEST RN 523977-56-2 DICTIONARY FILE UPDATES: 1 JUN 2003 HIGHEST RN 523977-56-2

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> e fumitremorgin

| E1 | 4 | FUMITOXIN/BI |
|-----|-----|--------------------|
| E2 | 1 | FUMITREMORGEN/BI |
| E3 | 36> | FUMITREMORGIN/BI |
| E4 | 3 | FUMJUDAINE/BI |
| E5 | 1 | FUMMITE/BI |
| E6 | 2 | FUMOFICIN/BI |
| E7 | 1. | FUMOFICINAL/BI |
| E8 | 1 | FUMOFICINALINE/BI |
| E9 | 1 | FUMOFICINAMINE/BI |
| E10 | 145 | FUMONISIN/BI |
| E11 | 7 | FUMOSA/BI |
| E12 | 1 | FUMOSIAVELLANEA/BI |

```
=> s e1-e3
```

4 FUMITOXIN/BI

1 FUMITREMORGEN/BI

36 FUMITREMORGIN/BI

L1 40 (FUMITOXIN/BI OR FUMITREMORGEN/BI OR FUMITREMORGIN/BI)

=> d 11 1-40

L1 ANSWER 1 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 119066-64-7 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14-octahydro-9-methoxy-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,12.alpha.,14a.beta.)]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 12.beta.-Fumitremorgin C

FS STEREOSEARCH

MF C22 H25 N3 O3

SR CA

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, TOXCENTER (*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1957 TO DATE)

2 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 2 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 118974-02-0 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-9-methoxy-12-(2-methyl-1-propenyl)-, (5aS,12S,14aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14dione, 1,2,3,5a,6,11,12,14a-octahydro-9-methoxy-12-(2-methyl-1-propenyl)-,
[5aS-(5a.alpha.,12.beta.,14a.alpha.)]-

OTHER NAMES:

CN 12.alpha.-Fumitremorgin C

CN Fumitremorgin C

FS STEREOSEARCH

MF C22 H25 N3 O3

SR CA

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.

- 28 REFERENCES IN FILE CA (1957 TO DATE)
 - 1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 28 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 3 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 111768-16-2 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-12-(2-methyl-1-propenyl)-, (5aS,12S,14aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-12-(2-methyl-1-propenyl)-, [5aS-(5a.alpha.,12.beta.,14a.alpha.)]-

OTHER NAMES:

CN (+)-Demethoxyfumitremorgin C

CN Demethoxyfumitremorgin C

FS STEREOSEARCH

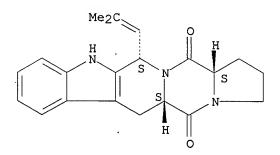
MF C21 H23 N3 O2

SR CA

LC STN Files: BEILSTEIN*, BIOSIS, CA, CAPLUS, CASREACT, CHEMINFORMRX, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 16 REFERENCES IN FILE CA (1957 TO DATE)
- 16 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 4 OF 40 REGISTRY COPYRIGHT 2003 ACS

```
RN
     111427-99-7 REGISTRY
CN
     5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-
     dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-12-(2-
     methyl-1-propenyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]- (9CI)
     (CA INDEX NAME)
OTHER NAMES:
     12,13-Dihydroxyfumitremorgin C
CN
CN
     TR 3
     TR 3 toxin
CN
CN
     Verruculogen TR 3
FS
     STEREOSEARCH
MF
     C22 H25 N3 O5
SR
LC
     STN Files:
                  BEILSTEIN*, CA, CAPLUS, CASREACT, TOXCENTER
         (*File contains numerically searchable property data)
```

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

7 REFERENCES IN FILE CA (1957 TO DATE) 7 REFERENCES IN FILE CAPLUS (1957 TO DATE) L1 ANSWER 5 OF 40 REGISTRY COPYRIGHT 2003 ACS RN 111427-98-6 REGISTRY CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-11-(3-methyl-2butenyl)-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a. alpha.)]- (9CI) (CA INDEX NAME) OTHER NAMES: CN Demethoxyfumitremorgin B FS STEREOSEARCH MF C26 H31 N3 O4 SR

BEILSTEIN*, CA, CAPLUS, CASREACT, TOXCENTER

(*File contains numerically searchable property data)

Absolute stereochemistry.

LC

- 4 REFERENCES IN FILE CA (1957 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 6 OF 40 REGISTRY COPYRIGHT 2003 ACS ·

RN 111080-12-7 REGISTRY

CN 1H,5H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,6,14(5aH,14aH)-trione, 2,3,11,12-tetrahydro-5a-hydroxy-9-methoxy-11-(3-methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, (5aR,12S,14aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 1H,5H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole5,6,14(5aH,14aH)-trione, 2,3,11,12-tetrahydro-5a-hydroxy-9-methoxy-11-(3methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, [5aR(5a.alpha.,12.beta.,14a.alpha.)]-

OTHER NAMES:

CN 13-Oxofumitremorgin B

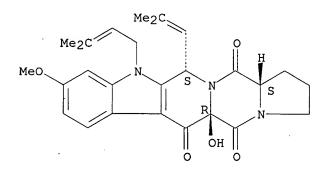
FS STEREOSEARCH

MF C27 H31 N3 O5

SR CA

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, TOXCENTER (*File contains numerically searchable property data)

Absolute stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1957 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 7 OF 40 REGISTRY COPYRIGHT 2003 ACS

```
RN
     107977-03-7 REGISTRY
CN
     5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-
     dione, 1,2,3,5a,6,11,12,14a-octahydro-6,9-dimethoxy-11-(3-methyl-2-
     butenyl)-12-(2-methyl-1-propenyl)-, (5a.alpha.,6.beta.,12.alpha.,14a.beta.
     )- (9CI)
              (CA INDEX NAME)
OTHER CA INDEX NAMES:
     5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-
CN
     dione, 1,2,3,5a,6,11,12,14a-octahydro-6,9-dimethoxy-11-(3-methyl-2-
     butenyl)-12-(2-methyl-1-propenyl)-, (5a.alpha.,6.beta.,12.alpha.,14a.beta.
     )-(.+-.)-
OTHER NAMES:
CN
     (.+-.)-12-Deoxy-13-O-methyl-12-epifumitremorgin B
     STEREOSEARCH
FS
MF
     C28 H35 N3 O4
SR
     CA
LC
     STN Files:
                  BEILSTEIN*, CA, CAPLUS, CASREACT
         (*File contains numerically searchable property data)
Relative stereochemistry.
              Me<sub>2</sub>C
```

1 REFERENCES IN FILE CA (1957 TO DATE)
1 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 8 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 107977-02-6 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-6-hydroxy-9-methoxy-11-(3-methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, (5a.alpha.,6.beta.,12.alpha.,14a.beta.)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14dione, 1,2,3,5a,6,11,12,14a-octahydro-6-hydroxy-9-methoxy-11-(3-methyl-2butenyl)-12-(2-methyl-1-propenyl)-, (5a.alpha.,6.beta.,12.alpha.,14a.beta.
)-(.+-.)-

OTHER NAMES:

CN (.+-.)-12-Deoxy-12-epifumitremorgin B

FS STEREOSEARCH

MF C27 H33 N3 O4

SR CA

LC STN Files: BEILSTEIN*, BIOSIS, CA, CAPLUS, CASREACT, TOXCENTER (*File contains numerically searchable property data)

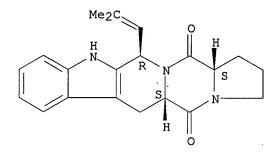
Relative stereochemistry.

- 1 REFERENCES IN FILE CA (1957 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 9 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 106292-68-6 REGISTRY
- CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-12-(2-methyl-1-propenyl)-, [5aS-(5a.alpha.,12.alpha.,14a.alpha.)]- (9CI) (CA INDEX NAME)

OTHER NAMES:

- CN Demethoxy-3-epifumitremorgin C
- FS STEREOSEARCH
- MF C21 H23 N3 O2
- SR CA
- LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMINFORMRX (*File contains numerically searchable property data)

Absolute stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1957 TO DATE)
- . 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 10 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 106211-91-0 REGISTRY
- CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,12.alpha.,14a.beta.)]- (9CI) (CA INDEX NAME)
- OTHER NAMES:
- CN Demethoxy-12-epifumitremorgin C
- FS STEREOSEARCH

```
MF C21 H23 N3 O2
```

SR CA

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMINFORMRX (*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1957 TO DATE)

1 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 11 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 80558-95-8 REGISTRY

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, [9'.alpha.(S*),9'a.beta.]- (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, [9'.alpha.(S*),9'a.beta.]-(.+-.)-

OTHER NAMES:

CN (.+-.)-FTG

CN (.+-.)-Fumitremorgin G

CN (.+-.)-Tryptoquivaline G

FS STEREOSEARCH

MF C23 H20 N4 O5

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

Relative stereochemistry.

- 2 REFERENCES IN FILE CA (1957 TO DATE)
 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 12 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 71658-19-0 REGISTRY

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
1'-(acetyloxy)-1',3,4,9'a-tetrahydro-4-[2-(1-hydroxy-2-methylpropyl)-4-oxo-3(4H)-quinazolinyl]-2'-methyl-, [2'S-[2'.alpha.,9'.beta.[4R*(R*)],9'a.alpha.]]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Deacetyl-12-epifumitremorgin D monoacetate

CN Epideacetyl-FTD monoacetate

FS STEREOSEARCH

MF C28 H28 N4 O7

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- · 1 REFERENCES IN FILE CA (1957 TO DATE)
 - 1 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 13 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 71658-06-5 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1'-(acetyloxy)-4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-tetrahydro-2'-methyl-, [2'S-[2'.alpha.,9'.beta.[4R*(R*)],9'a.alpha.]]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN FTM acetate

CN Fumitremorgin M acetate

CN Tryptoquivaline M acetate

MF C30 H30 N4 O8

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

- 1 REFERENCES IN FILE CA (1957 TO DATE).
- 1 REFERENCES IN FILE CAPLUS (1957 TO DATE)

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L1 ANSWER 14 OF 40 REGISTRY COPYRIGHT 2003 ACS
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RN 69575-59-3 REGISTRY

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
4-[2-[(1S)-1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-, (2S,2'S,4S,9'aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-, [2'S-[2'.alpha.,9'.beta.[4R*(R*)],9'a.alpha.]]-

OTHER NAMES:

CN FTM

CN Fumitremorgin M

CN Tryptoquivaline M

FS STEREOSEARCH

MF C28 H28 N4 O7

LC STN Files: BEILSTEIN*, CA, CAPLUS, NAPRALERT (*File contains numerically searchable property data)

Absolute stereochemistry.

- 2 REFERENCES IN FILE CA (1957 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 15 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 69483-20-1 REGISTRY

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, (2S,4S,9'aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, [9'S-[9'.alpha.(R*),9'a.beta.]]-

OTHER NAMES:

CN 19-Epifumitremorgin G

CN 19-Epitryptoquivaline G

CN FTL

CN Fumitremorgin L

CN Tryptoquivaline L

FS STEREOSEARCH

MF C23 H20 N4 O5

LC STN Files: BEILSTEIN*, CA, CAPLUS, NAPRALERT

(*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 4 REFERENCES IN FILE CA (1957 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 16 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 68817-01-6 REGISTRY

CN Fumitoxin D (9CI) (CA INDEX NAME)

MF Unspecified

CI MAN

LC STN Files: CA, CAPLUS, TOXCENTER

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

- 2 REFERENCES IN FILE CA (1957 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 17 OF 40 REGISTRY COPYRIGHT 2003 ACS

```
CN
     Fumitoxin C (9CI) (CA INDEX NAME)
MF
     Unspecified
CI
     MAN
LC
                  CA, CAPLUS, TOXCENTER
     STN Files:
    STRUCTURE DIAGRAM IS NOT AVAILABLE ***
               2 REFERENCES IN FILE CA (1957 TO DATE)
               2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
L1
     ANSWER 18 OF 40 REGISTRY COPYRIGHT 2003 ACS
RN
     68816-99-9 REGISTRY
CN
     Fumitoxin B (9CI) (CA INDEX NAME)
MF
     Unspecified
CI
     MAN
     STN Files:
LC
                  CA, CAPLUS, TOXCENTER
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
               2 REFERENCES IN FILE CA (1957 TO DATE)
               2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
L1
     ANSWER 19 OF 40 REGISTRY COPYRIGHT 2003 ACS
RN
     66419-35-0 REGISTRY
CN
     Fumitoxin A (9CI) (CA INDEX NAME)
MF
     Unspecified
CI
     MAN
LC
     STN Files:
                  BIOSIS, CA, CAPLUS, TOXCENTER
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
               9 REFERENCES IN FILE CA (1957 TO DATE)
               9 REFERENCES IN FILE CAPLUS (1957 TO DATE)
     ANSWER 20 OF 40 REGISTRY COPYRIGHT 2003 ACS
L1
RN
     66212-52-0 REGISTRY
CN
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     1'-acetyl-1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-,
     [2'S-[2'.alpha.,9'.beta.(S*),9'a.alpha.]]- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     FTJ acetate
     N-Acetylfumitremorgin J
CN
CN
     N-Acetyltryptoquivaline J
MF
     C24 H20 N4 O5
LC
     STN Files:
                BEILSTEIN*, CA, CAPLUS
         (*File contains numerically searchable property data)
```

RN

68817-00-5 REGISTRY

- 1 REFERENCES IN FILE CA (1957 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 21 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 66212-51-9 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, (2S,2'S,4R,9'aR)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, [2'S-[2'.alpha.,9'.beta.(S*),9'a.alpha.]]-

OTHER NAMES:

CN FTJ

CN Fumitremorgin J

CN Tryptoquivaline J

FS STEREOSEARCH

MF C22 H18 N4 O4

LC STN Files: ANABSTR, BEILSTEIN*, CA, CAPLUS, NAPRALERT, TOXCENTER (*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).

- **PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
 - 3 REFERENCES IN FILE CA (1957 TO DATE)
 - 3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 22 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 66180-23-2 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-[2-(2-methyl-1-oxopropyl)-4-oxo-3(4H)-quinazolinyl]-, (2S,4R,9'aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-[2-(2-methyl-1-oxopropyl)-4-oxo-3(4H)-quinazolinyl]-, [9'S-[9'.alpha.(S*),9'a.beta.]]-OTHER NAMES:
- CN FTI
- CN Fumitremorgin I
- CN Tryptoquivaline I

MF C27 H26 N4 O6

LC STN Files: BEILSTEIN*, CA, CAPLUS, NAPRALERT (*File contains numerically searchable property data)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1957 TO DATE)

3 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 23 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 61949-68-6 REGISTRY

CN Spiro[furan-2(5H);9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1'-(acetyloxy)-1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, [2'S-[2'.alpha.,9'.beta.(S*),9'a.alpha.]]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN FTE acetate

CN Fumitremorgin E acetate

CN Tryptoquivaline E acetate

MF C24 H20 N4 O6

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

```
2 REFERENCES IN FILE CA (1957 TO DATE)
2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
```

L1 ANSWER 24 OF 40 REGISTRY COPYRIGHT 2003 ACS RN 61949-67-5 REGISTRY CN Spiro[furan-2(5H), 9'-[9H]imidazo[1, 2-a]indole]-3', 5(2'H)-dione,1', 3, 4, 9'a-tetrahydro-1'-hydroxy-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-,(2S,2'S,4S,9'aS)- (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, [2'S-[2'.alpha.,9'.beta.(R*),9'a.alpha.]]-OTHER NAMES: CN FTH CN Fumitremorgin H CN Tryptoquivaline H FS STEREOSEARCH MF C22 H18 N4 O5 LC STN Files: BEILSTEIN*, CA, CAPLUS, NAPRALERT

(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1957 TO DATE)

3 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 25 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 61897-92-5 REGISTRY

CN FTG acetate

CN Fumitremorgin G acetate

CN Tryptoquivaline G acetate

MF C25 H22 N4 O6

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

2 REFERENCES IN FILE CA (1957 TO DATE)
2 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 26 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 61897-91-4 REGISTRY

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, (2S,4R,9'aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-4-(4-oxo-3(4H)-quinazolinyl)-, [9'S-[9'.alpha.(S*),9'a.beta.]]-

OTHER NAMES:

CN FTG

CN Fumitremorgin G

CN Tryptoquivaline G

FS STEREOSEARCH

MF C23 H20 N4 O5

LC STN Files: AGRICOLA, BEILSTEIN*, BIOSIS, CA, CAPLUS, NAPRALERT, TOXCENTER

(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

```
6 REFERENCES IN FILE CAPLUS (1957 TO DATE)
     ANSWER 27 OF 40 REGISTRY COPYRIGHT 2003 ACS
L1
RN
     61897-90-3 REGISTRY
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
CN
     1'-acetyl-1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-,
     [2'S-[2'.alpha.,9'.beta.(R*),9'a.alpha.]]- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     FTF acetate
CN
     Fumitremorgin F acetate
CN
    N-Acetylfumitremorgin F
CN
     N-Acetyltryptoquivaline F
MF
     C24 H20 N4 O5
                  BEILSTEIN*, CA, CAPLUS .
LC ·
     STN Files:
         (*File contains numerically searchable property data)
```

6 REFERENCES IN FILE CA (1957 TO DATE)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

```
2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
     ANSWER 28 OF 40 REGISTRY COPYRIGHT 2003 ACS
L1
RN
     61897-89-0 REGISTRY
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
CN
     1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, (2S,2'S,4S,9'aR)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     1', 3, 4, 9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-,
     [2'S-[2'.alpha.,9'.beta.(R*),9'a.alpha.]]-
OTHER NAMES:
CN
     FTF
CN
     Fumitremorgin F
CN
     Tryptoquivaline F
FS
     STEREOSEARCH
MF
     C22 H18 N4 O4
LC
     STN Files:
                   ANABSTR, BEILSTEIN*, CA, CAPLUS, NAPRALERT, TOXCENTER
```

(*File contains numerically searchable property data)

2 REFERENCES IN FILE CA (1957 TO DATE)

Absolute stereochemistry. Rotation (-).

- 4 REFERENCES IN FILE CA (1957 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 29 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 61897-88-9 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1'-(acetyloxy)-1',3,4,9'a-tetrahydro-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, [2'S-[2'.alpha.,9'.beta.(R*),9'a.alpha.]]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN FTH acetate

CN Fumitremorgin H acetate

CN Tryptoquivaline H acetate

MF C24 H20 N4 O6

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1957 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 30 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 61897-87-8 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-, (2S,2'S,4R,9'aS)- (9CI) (CA INDEX NAME)

```
OTHER CA INDEX NAMES:
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-4-(4-oxo-3(4H)-quinazolinyl)-,
     [2'S-[2'.alpha.,9'.beta.(S*),9'a.alpha.]]-
OTHER NAMES:
    FTE
CN
CN
     Fumitremorgin E
CN
     Tryptoquivaline E
FS
     STEREOSEARCH
MF
     C22 H18 N4 O5
LC
     STN Files:
                  AGRICOLA, BEILSTEIN*, BIOSIS, CA, CAPLUS, NAPRALERT,
       TOXCENTER
         (*File contains numerically searchable property data)
```

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

```
4 REFERENCES IN FILE CA (1957 TO DATE)
               4 REFERENCES IN FILE CAPLUS (1957 TO DATE)
L1
     ANSWER 31 OF 40 REGISTRY COPYRIGHT 2003 ACS
RN
     60676-61-1 REGISTRY
CN
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     1',3,4,9'a-tetrahydro-2'-methyl-4-[2-(2-methyl-1-oxopropyl)-4-oxo-3(4H)-
     quinazolinyl]-, (2S,2'S,4R,9'aR)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     1',3,4,9'a-tetrahydro-2'-methyl-4-[2-(2-methyl-1-oxopropyl)-4-oxo-3(4H)-
     quinazolinyl]-, [2'S-[2'.alpha.,9'.beta.(S*),9'a.alpha.]]-
OTHER NAMES:
CN
     1'-Deoxytryptoquivalone
CN
     Deoxynortryptoquivalone
```

Deoxytryptoquivaline B CN Fumitremorgin N

CN Tryptoquivaline N

MF C26 H24 N4 O5

CN

LC STN Files: BEILSTEIN*, BIOSIS, CA, CAPLUS, NAPRALERT, TOXCENTER (*File contains numerically searchable property data)

- 3 REFERENCES IN FILE CA (1957 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 32 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 60676-59-7 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
 1'-(acetyloxy)-4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)quinazolinyl]-1',3,4,9'a-tetrahydro-2'-methyl-, [2'S-

[2'.alpha.,9'.beta.[4S*(R*)],9'a.alpha.]]- (9CI) (CA INDEX NAME) OTHER NAMES:

CN FTD acetate

CN Fumitremorgin D acetate

CN Norisotryptoquivaline acetate

CN Nortryptoquivaline acetate

CN O-Acetyl-2'-demethyltryptoquivaline A

DR 61949-66-4, 71658-07-6

MF C30 H30 N4 O8

LC STN Files: BEILSTEIN*, CA, CAPLUS, TOXCENTER

(*File contains numerically searchable property data)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1957 TO DATE)

1 REFERENCES IN FILE CAPLUS (1957 TO DATE)

```
ANSWER 33 OF 40 REGISTRY COPYRIGHT 2003 ACS
L1
RN
     60676-56-4 REGISTRY
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
CN
     4-[2-[(1S)-1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-
     1',3,4,9'a-tetrahydro-1'-hydroxy-2'-methyl-, (2S,2'S,4R,9'aS)- (9CI)
     INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-
     tetrahydro-1'-hydroxy-2'-methyl-, [2'S-[2'.alpha.,9'.beta.[4S*(R*)],9'a.al
     pha.]]-
OTHER NAMES:
     2'-Demethyltryptoquivaline
CN
CN
     Fumitremorgin D
CN
CN
     Nortryptoquivaline
     Nortryptoquivaline A
CN
CN
     Tryptoquivaline D
FS
     STEREOSEARCH
MF
     C28 H28 N4 O7
LC
     STN Files:
                  AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS,
       NAPRALERT, TOXCENTER
         (*File contains numerically searchable property data)
```

Absolute stereochemistry. Rotation (+).

- **PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
 - 4 REFERENCES IN FILE CA (1957 TO DATE)
 - 1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
 - 4 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 34 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 55387-47-8 REGISTRY
- CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione, 1'-(acetyloxy)-4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-tetrahydro-2',2'-dimethyl-, [9'S-[9'.alpha.[4S*(R*)],9'a.beta.]]- (9CI) (CA INDEX NAME)
- OTHER NAMES:
- CN Fumitremorgin C acetate
- CN Tryptoquivaline A acetate

```
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
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2 REFERENCES IN FILE CA (1957 TO DATE)
2 REFERENCES IN FILE CAPLUS (1957 TO DATE)
```

```
L1
     ANSWER 35 OF 40 REGISTRY COPYRIGHT 2003 ACS
RN
     55387-45-6 REGISTRY
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
CN
     4-[2-[(1S)-1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-
     1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-, (2S,4R,9'aS)- (9CI)
     INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
     4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-
     tetrahydro-1'-hydroxy-2',2'-dimethyl-, [9'S-[9'.alpha.[4S*(R*)],9'a.beta.]
     ] -
OTHER NAMES:
     Fumitremorgin C
CN
     Tryptoquivaline
CN
     Tryptoquivaline A
CN
     Tryptoquivaline C
FS
     STEREOSEARCH
MF
     C29 H30 N4 O7
LC
     STN Files:
                 AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA,
       CANCERLIT, CAPLUS, EMBASE, MEDLINE, NAPRALERT, TOXCENTER
         (*File contains numerically searchable property data)
```

Absolute stereochemistry. Rotation (+).

12 REFERENCES IN FILE CA (1957 TO DATE)

12 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 36 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 54009-33-5 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methylbutyl)-12-(2-methylpropyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Tetrahydrofumitremorgin B

CN Tetrahydrolanosulin

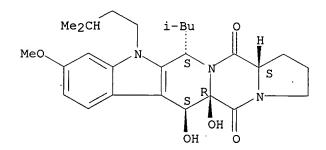
FS STEREOSEARCH

MF C27 H37 N3 O5

LC STN Files: BEILSTEIN*, CA, CAPLUS, TOXCENTER

(*File contains numerically searchable property data)

Absolute stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1957 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 37 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 54009-32-4 REGISTRY
- CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methylbutyl)-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]- (9CI) (CA INDEX NAME)

```
OTHER NAMES:
     22,23-Dihydrofumitremorgin B
     Dihydrofumitremorgin B
CN
      Dihydrolanosulin.
     STEREOSEARCH
FS
MF
      C27 H35 N3 O5
                    BEILSTEIN*, CA, CAPLUS, TOXCENTER
LC
      STN Files:
          (*File contains numerically searchable property data)
Absolute stereochemistry.
               Me<sub>2</sub>C
     Me<sub>2</sub>CH
```

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**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
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3 REFERENCES IN FILE CA (1957 TO DATE)
3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
```

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L1 ANSWER 38 OF 40 REGISTRY COPYRIGHT 2003 ACS
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RN 12626-18-5 REGISTRY

CN 5H,12H-3,4-Dioxa-5a,11a,15a-triazacyclooct[lm]indeno[5,6-b]fluorene-11,15(2H,13H)-dione, 1,10,10a,14,14a,15b-hexahydro-10a-hydroxy-7-methoxy-2,2-dimethyl-10-[(3-methyl-2-butenyl)oxy]-5-(2-methyl-1-propenyl)-, (5R,10S,10aR,14aS,15bS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,12H-3,4-Dioxa-5a,11a,15a-triazacyclooct[lm]indeno[5,6-b]fluorene-11,15(2H,13H)-dione, 1,10,10a,14,14a,15b-hexahydro-10a-hydroxy-7-methoxy-2,2-dimethyl-10-[(3-methyl-2-butenyl)oxy]-5-(2-methyl-1-propenyl)-, [5R-(5.alpha.,10.alpha.,10a.alpha.,14a.alpha.,15b.alpha.)]-

OTHER NAMES: CN Fumitremorgen A

CN Fumitremorgin A

FS STEREOSEARCH

MF C32 H41 N3 O7

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAPLUS, DDFU, DRUGU, EMBASE, MEDLINE, NAPRALERT, RTECS*, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.

24 REFERENCES IN FILE CA (1957 TO DATE)

24 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 39 OF 40 REGISTRY COPYRIGHT 2003 ACS

RN 12626-17-4 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, (5aR,6S,12S,14aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]-

OTHER NAMES:

CN Fumitremorgin B

CN Lanosulin

CN NA 209B

FS STEREOSEARCH

DR 40451-43-2

MF C27 H33 N3 O5

CI COM

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAPLUS, CASREACT, DDFU, DRUGU, EMBASE, MEDLINE, NAPRALERT, RTECS*, SPECINFO, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.

- 42 REFERENCES IN FILE CA (1957 TO DATE)
- 1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 42 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 40 OF 40 REGISTRY COPYRIGHT 2003 ACS
- RN 11100-25-7 REGISTRY
- CN Fumitremorgin (9CI) (CA INDEX NAME)
- MF Unspecified
- CI MAN
- LC STN Files: AGRICOLA, BIOBUSINESS, BIOSIS, CA, CAPLUS, TOXCENTER
- *** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
 - 9 REFERENCES IN FILE CA (1957 TO DATE)
 - 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
 - 9 REFERENCES IN FILE CAPLUS (1957 TO DATE)

=> file caplus
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 83.06 83.27

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 13:41:20 ON 03 JUN 2003
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FILE COVERS 1907 - 3 Jun 2003 VOL 138 ISS 23 FILE LAST UPDATED: 2 Jun 2003 (20030602/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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L2 124 L1

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=> s hiv or herpes or viral or virus

49538 HIV

21375 HERPES

119994 VIRAL

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L3 315130 HIV OR HERPES OR VIRAL OR VIRUS

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     Application of a human multidrug transporter (abcg2) variant as selectable
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     marker in gene transfer to progenitor cells and in gene therapy
ΙN
     Nemet, Katalin; Varadi, Gyorgy; Cervenak, Judit; Ujhelly, Olga; Sarkadi,
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PA
     Solvo Biotechnology Inc., Hung.
SO
     PCT Int. Appl., 44 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
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FAN.CNT 2
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                       Α
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     ANSWER 2 OF 3 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:46901 CAPLUS
DN
     137:125308
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     Solid phase synthesis of fumitremorgin-type and other indole alkaloids
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ΑU
     van Loevezijn, Arnold; Rodenko, Boris; Sorm, Willem P.; van Maarseveen,
     Jan H.; Stegman, Karel; Visser, Geb M.; van Delft, Floris L.; Koomen,
CS
     Laboratory of Organic Chemistry, Institute for Molecular Chemistry,
     University of Amsterdam, Amsterdam, NL-1018 WS, Neth.
SO
     Innovation and Perspectives in Solid Phase Synthesis & Combinatorial
     Libraries: Peptides, Proteins and Nucleic Acids--Small Molecule Organic
     Chemistry Diversity, Collected Papers, International Symposium, 6th, York,
     United Kingdom, Aug. 31-Sept. 4, 1999 (2001), Meeting Date 1999, 367-370.
     Editor(s): Epton, Roger. Publisher: Mayflower Scientific Ltd.,
     Kingswinford, UK.
     CODEN: 69CEGV; ISBN: 0-9515735-3-5
DT
     Conference
LA
     English
RE.CNT 6
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              ALL CITATIONS AVAILABLE IN THE RE FORMAT
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     ANSWER 3 OF 3 CAPLUS COPYRIGHT 2003 ACS
AN
     1999:511178 CAPLUS
DN
     131:143080
ΤT
     A multidrug resistance protein associated with antitumor drug resistance
     in breast cancer and a cDNA encoding it
IN·
     Ross, Douglas D.; Doyle, L. Austin; Abruzzo, Lynne
PA
     University of Maryland, Baltimore, USA
     PCT Int. Appl., 79 pp.
SO
     CODEN: PIXXD2
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· IE, FI US 6313277 B1 20011106 US 1999-245808 JP 2002502592 T2 20020129 JP 2000-530538 US 2003036645 A1 20030220 US 2001-961086 19980205 PRAI US 1998-73763P P US 1999-245808 A3 19990205 WO 1999-US2577 W 19990205 RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT => d his (FILE 'HOME' ENTERED AT 13:36:13 ON 03 JUN 2003) FILE 'REGISTRY' ENTERED AT 13:36:28 ON 03 JUN 2003 E FUMITREMORGIN L1 40 S E1-E3 FILE 'CAPLUS' ENTERED AT 13:41:20 ON 03 JUN 2003 L2124 S L1 L3 315130 S HIV OR HERPES OR VIRAL OR VIRUS 0 S L2 AND L3 E CHEMOTHERAPUTIC 35581 S E1-E4 3 S L2 AND L5 L6 =>

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1994:94935 CAPLUS ΑN 120:94935 ĎΝ ΤI Cross-resistance to diverse drugs is associated with primary cisplatin resistance in ovarian cancer cell lines AU Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter J.; Ozols, Robert F.; Hamilton, Thomas C. Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA CS Cancer Research (1993), 53(21), 5225-32 SO CODEN: CNREA8; ISSN: 0008-5472 DTJournal LΑ English CC 1-6 (Pharmacology) AB The authors have previously obtained, by exposure to near continuous increasing concns. of cisplatin, a panel of human ovarian cancer cell lines that exhibit a wide range of primary resistance to the drug (9- to >400-fold). These cells had strikingly increased (4- to 50-fold) levels of glutathione (GSH) as compared with the drug-sensitive cells of origin (A. K. Godwin et al., Proc. Natl. Acad. Sci. USA, 89: 3070-3074, 1992). Using this panel of resistant cell lines, the authors evaluated cross-resistance to classical alkylating agents, natural product drugs, and irradn. Cross-resistance to carboplatin paralleled that of cisplatin, culminating is .apprx. 250-fold Similarly, melphalan cross-resistance resistance. continued to increase to >400-fold and again paralleled the primary cisplatin resistance. Cell lines with low to very high levels of resistance to cisplatin are 8-850-fold resistant to the epipodophyllotoxin deriv. etoposide. Cross-resistance is also obsd. for other natural product drugs, including Adriamycin (.apprx.80-fold), mitoxantrone (.apprx.440-fold), and taxol (.apprx.40-fold). Cross-resistance to irradn. is, however, modest (<2-fold). The cells with greatest primary resistance to cisplatin most commonly had the highest cross-resistance to the other drugs examd. The cross-resistance to the natural produce category drugs was found not to be mediated by the products of either the multidrug resistance 1 (MDR1) or multidrug resistance -assocd. protein (MRP) genes based on lack of coordinate increased expression or amplification of these genes as assessed by Northern and Southern blot analyses. Also, verapamil failed to markedly increase drug sensitivity. Although there was no indication that these natural product drug efflux pumps were operative, the authors obsd. decreased doxorubicin accumulation in these cell lines cross-resistant to natural products. Alternations in DNA topoisomerase II mRNA levels, which were obsd. in human tumor cell lines selected in vitro for resistance to etoposide or teniposide, were not detected. Only intracellular levels of GSH correlated with cross-resistance to these diverse anticancer agents and partial loss of resistance was assocd. with a marked decrease in glutathione levels. In the absence of alternative mechanisms, the authors speculate that the very broad clin. relevant cross-resistance seen in this model system may, at least in part, be the direct result of GSH-mediated drug inactivation or may be due to a combination of GSH conjugation to drug and conjugate efflux mediated by the putative ATP-dependent glutathione S-conjugate export pump. STcisplatin resistance neoplasm cross resistance glutathione ΙT Neoplasm inhibitors (cisplatin as, resistance to, cross-resistance to,

in humans cells, GSH in mechanism of)
IT Radiation
 (cross-resistance of, in neoplasm of humans, to cisplatin
 resistance, GSH in mechanism of)
IT Biological transport

(of doxorubicin, in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in) IT Drug resistance (to cisplatin, cross-resistance to, in neoplasm of humans, GSH in mechanism of) IT 148-82-3, Melphalan 33069-62-4, Taxol 33419-42-0, VP-16 41575-94-4, Carboplatin 65271-80-9, Mitoxantrone RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of) ΙT 23214-92-8, Adriamycin RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of, transport in relation to) IT 70-18-8, Glutathione, biological studies RL: BIOL (Biological study) (in neoplasm of humans resistance to cisplatin, crossresistance to other agents in) IT **15663-27-1**, Cisplatin RL: BIOL (Biological study)

(resistance to, in neoplasm of humans, cross-

resistance in, GSH in mechanism of)

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1994:94935 CAPLUS AN DN 120:94935 ΤI Cross-resistance to diverse drugs is associated with primary cisplatin resistance in ovarian cancer cell lines ΑU Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter J.; Ozols, Robert F.; Hamilton, Thomas C. Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA CS SO Cancer Research (1993), 53(21), 5225-32 CODEN: CNREA8; ISSN: 0008-5472 DTJournal LΑ English CC 1-6 (Pharmacology) AB The authors have previously obtained, by exposure to near continuous increasing concns. of cisplatin, a panel of human ovarian cancer cell lines that exhibit a wide range of primary resistance to the drug (9- to >400-fold). These cells had strikingly increased (4- to 50-fold) levels of glutathione (GSH) as compared with the drug-sensitive cells of origin (A. K. Godwin et al., Proc. Natl. Acad. Sci. USA, 89: 3070-3074, 1992). Using this panel of resistant cell lines, the authors evaluated cross-resistance to classical alkylating agents, natural product drugs, and irradn. Cross-resistance to carboplatin paralleled that of cisplatin, culminating is .apprx. 250-fold resistance. Similarly, melphalan cross-resistance continued to increase to >400-fold and again paralleled the primary cisplatin resistance. Cell lines with low to very high levels of resistance to cisplatin are 8-850-fold resistant to the epipodophyllotoxin deriv. etoposide. Cross-resistance is also obsd. for other natural product drugs, including Adriamycin (.apprx.80-fold), mitoxantrone (.apprx.440-fold), and taxol (.apprx.40-fold). Cross-resistance to irradn. is, however, modest (<2-fold). The cells with greatest primary resistance to cisplatin most commonly had the highest cross-resistance to the other drugs examd. The cross-resistance to the natural produce category drugs was found not to be mediated by the products of either the multidrug resistance 1 (MDR1) or multidrug resistance -assocd. protein (MRP) genes based on lack of coordinate increased expression or amplification of these genes as assessed by Northern and Southern blot analyses. Also, verapamil failed to markedly increase drug sensitivity. Although there was no indication that these natural product drug efflux pumps were operative, the authors obsd. decreased doxorubicin accumulation in these cell lines cross-resistant to natural products. Alternations in DNA topoisomerase II mRNA levels, which were obsd. in human tumor cell lines selected in vitro for resistance to etoposide or teniposide, were not detected. Only intracellular levels of GSH correlated with cross-resistance to these diverse anticancer agents and partial loss of resistance was assocd. with a marked decrease in glutathione levels. In the absence of alternative mechanisms, the authors speculate that the very broad clin. relevant cross-resistance seen in this model system may, at least in part, be the direct result of GSH-mediated drug inactivation or may be due to a combination of GSH conjugation to drug and conjugate efflux mediated by the putative ATP-dependent glutathione S-conjugate export pump. stcisplatin resistance neoplasm cross resistance glutathione Neoplasm inhibitors (cisplatin as, resistance to, cross-resistance to, in humans cells, GSH in mechanism of) IT Radiation (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of) IT Biological transport

(of doxorubicin, in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in) IT Drug resistance (to cisplatin, cross-resistance to, in neoplasm of humans, GSH in mechanism of) ΙT 148-82-3, Melphalan 33069-62-4, Taxol 33419-42-0, VP-16 41575-94-4, 65271-80-9, Mitoxantrone Carboplatin RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of) ΙT 23214-92-8, Adriamycin RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of, transport in relation to) ΙT 70-18-8, Glutathione, biological studies RL: BIOL (Biological study) (in neoplasm of humans resistance to cisplatin, crossresistance to other agents in) IT**15663-27-1**, Cisplatin RL: BIOL (Biological study) (resistance to, in neoplasm of humans, cross-

resistance in, GSH in mechanism of)

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124:285652 DN ΤI Alterations in expression of the multidrug resistance-associated protein (MRP) gene in high-grade transitional cell carcinoma of the bladder ΑU Clifford, S. C.; Neal, D. E.; Lunec, J. CS Medical School, University of Newcastle-upon-Tyne, Newcastle-upon-Tyne, NE2 4HH, UK British Journal of Cancer (1996), 73(5), 659-66 SO CODEN: BJCAAI; ISSN: 0007-0920 PB Stockton DT Journal LΑ English CC 14-1 (Mammalian Pathological Biochemistry) Section cross-reference(s): 1 AB Expression of the MRP gene has been demonstrated in vitro to be a causal factor in non-P-glycoprotein-mediated multidrug resistance, and is implicated in resistance to a no. of the chemotherapeutic agents currently used in the treatment of high-grade transitional cell carcinoma (TCC) of the bladder (doxorubicin, epirubicin and vinblastine). Using a sensitive RT-PCR-based technique, we have quantified MRP mRNA levels in a series of untreated TCC (n=24), normal bladder (n=5) and control tissue and cell line samples. MRP mRNA was widely expressed and detectable in all samples analyzed, with considerable (up to 190-fold) variation obsd. between individual tumor samples. MRP mRNA levels found in TCC samples were lower than those detd. for normal peripheral mononucleocyte (2.3-fold) and testis (4.1-fold) samples, previously reported to be high-expressing tissues, and varied over a similar range to that obsd. in normal bladder samples. Results indicate that MRP mRNA levels in a greater proportion of high-grade (G3) bladder tumors (55%, 6/11) are significantly reduced (P=0.018) compared with low- and moderate-grade (G1/2) bladder tumors (8%, 1/13), and suggest that MRP mRNA levels frequently become reduced as a consequence of tumor progression to advanced, poorly differentiated disease. No correlation was apparent between MRP and MDR1 mRNA levels, thus providing no evidence to suggest common regulation of the two genes. In a limited no. of patients, no evidence was found to support a role for MRP mRNA levels as a determinant of response to chemotherapy in patients being uniformly treated with either cisplatin-methotrexate-vinblastine (n=6) or epirubicin-cisplatin-methotrexate (n=4) regimens. Similarly, no overall pattern of altered MRP mRNA expression was obsd. following chemotherapy in four patients from whom post chemotherapy biopsies were taken. This study provides a useful pilot investigation regarding the level, variation and pattern of MRP mRNA expression in TCC of the bladder, and suggests that further studies to establish the clin. significance of these variations are required. ST multidrug resistance protein gene bladder carcinoma ΙT Gene, animal RL: BSU (Biological study, unclassified); BIOL (Biological study) (MRP; multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder) ITProteins, specific or class RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(multidrug resistance-assocd. protein (MRP); multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder) Neoplasm inhibitors

(multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

ΙT Ribonucleic acids, messenger

IT

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

IT Bladder

(neoplasm, transitional cell carcinoma, multidrug resistance
-assocd. protein gene expression in human high-grade transitional cell
carcinoma of the bladder)

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

=>

ΑN 1996:96400 CAPLUS DN 124:193617 TI Non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line ΑU Naito, Seiji; Hasegawa, Shuji; Yokomizo, Akira; Koga, Hirofumi; Kotoh, Shuji; Kuwano, Michihiko; Kumazawa, Joichi CS Fac. Medicine, Kyushu Univ., Fukuoka, 812, Japan Japanese Journal of Cancer Research (1995), 86(11), 1112-18 SO CODEN: JJCREP; ISSN: 0910-5050 PB Japanese Cancer Association DT Journal LA English CC 1-6 (Pharmacology) AΒ A human bladder cancer cell line resistant to adriamycin (ADM), T24/ADM9 has been established in vitro by exposing T24 parent cells to progressively higher concns. of the drug over a period of 12 mo. T24/ADM9 cells were 9 times more resistant to ADM than the T24 parent, and showed various degrees of cross-resistance to an ADM deriv., vinca alkaloids and a DNA topoisomerase II (Topo II)-targeting agent, etoposide. No significant difference was obsd. in the cellular accumulation of ADM between the T24/ADM9 and T24 parent cells. A Northern blot anal. showed an overexpression of multidrug resistance -assocd. protein (MRP) mRNA, but no overexpression of multidrug resistance-1 (MDR1) mRNA was obsd. in the T24/ADM9 cells. A flow cytometric anal. showed that the MDR1 gene product, P-glycoprotein (Pgp), is not expressed on the T24/ADM9 cells. T24/ADM9 showed approx. the parental level of DNA Topo II catalytic activity. In Western blot and Northern blot analyses, however, the cellular level of DNA Topo II was apparently much lower in T24/ADM9 than in the T24 parent. Thus, these results suggest that a decreased cellular level of DNA Topo II and an overexpression of MRP gene may be responsible for the expression of an MDR phenotype in the T24/ADM9 cells and that such non-Pgp-mediated, atypical MDR may develop in bladder cancer treated with chemotherapy including ADM. ST atypical multidrug resistant bladder cancer cell Proteins, specific or class RL: BSU (Biological study, unclassified); BIOL (Biological study) (multidrug resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Gene, animal RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Neoplasm inhibitors (bladder carcinoma, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT Drug resistance (multi-, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) ΙT (neoplasm, carcinoma, inhibitors, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) ΙT 50-07-7, Mitomycin C 51-21-8, 5 Fluorouracil 57-22-7, Vincristine 865-21-4, Vinblastine **15663-27-1**, Cisplatin 33419-42-0, Etoposide 56420-45-2, Epirubicin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (cross-resistance; non-P-glycoprotein-mediated atypical

بنمنش ر ان

multidrug resistance in a human bladder cancer cell line)

142805-56-9, DNA topoisomerase II

RL: BOC (Biological occurrence); BSU (Biological study, unclassified);

BIOL (Biological study); OCCU (Occurrence)

(non-P-glycoprotein-mediated atypical multidrug resistance in

a human bladder cancer cell line in relation to decreased level of DNA

Topo II)

IT 25316-40-9, Adriamycin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

نسب ر آبيا

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(Uses)

(resistance to; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line)

124:193617 CAPLUS AN ΤI Non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line AU Naito, Seiji; Hasegawa, Shuji; Yokomizo, Akira; Koga, Hirofumi; Kotoh, Shuji; Kuwano, Michihiko; Kumazawa, Joichi CS Fac. Medicine, Kyushu Univ., Fukuoka, 812, Japan SO Japanese Journal of Cancer Research (1995), 86(11), 1112-18 CODEN: JJCREP; ISSN: 0910-5050 PB Japanese Cancer Association DTJournal LAEnglish CC 1-6 (Pharmacology) AB A human bladder cancer cell line resistant to adriamycin (ADM), T24/ADM9 has been established in vitro by exposing T24 parent cells to progressively higher concns. of the drug over a period of 12 mo. T24/ADM9 cells were 9 times more resistant to ADM than the T24 parent, and showed various degrees of cross-resistance to an ADM deriv., vinca alkaloids and a DNA topoisomerase II (Topo II)-targeting agent, etoposide. No significant difference was obsd. in the cellular accumulation of ADM between the T24/ADM9 and T24 parent cells. A Northern blot anal. showed an overexpression of multidrug resistance-assocd. protein (MRP) mRNA, but no overexpression of multidrug resistance-1 (MDR1) mRNA was obsd. in the T24/ADM9 cells. A flow cytometric anal. showed that the MDR1 gene product, P-glycoprotein (Pgp), is not expressed on the T24/ADM9 cells. T24/ADM9 showed approx. the parental level of DNA Topo II catalytic activity. In Western blot and Northern blot analyses, however, the cellular level of DNA Topo II was apparently much lower in T24/ADM9 than in the T24 parent. Thus, these results suggest that a decreased cellular level of DNA Topo II and an overexpression of MRP gene may be responsible for the expression of an MDR phenotype in the T24/ADM9 cells and that such non-Pgp-mediated, atypical MDR may develop in bladder cancer treated with chemotherapy including ADM. ST atypical multidrug resistant bladder cancer cell ΙT Proteins, specific or class RL: BSU (Biological study, unclassified); BIOL (Biological study) (multidrug resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Gene, animal RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Neoplasm inhibitors (bladder carcinoma, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT Drug resistance (multi-, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) ΙT Bladder (neoplasm, carcinoma, inhibitors, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) 50-07-7, Mitomycin C 51-21-8, 5 Fluorouracil 57-22-7, Vincristine 865-21-4, Vinblastine 15663-27-1, Cisplatin 33419-42-0, Etoposide 56420-45-2, Epirubicin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cross-resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT 142805-56-9, DNA topoisomerase II RL: BOC (Biological occurrence); BSU (Biological study, unclassified);

BIOL (Biological study); OCCU (Occurrence)

(non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to decreased level of DNA Topo II) 25316-40-9, Adriamycin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(resistance to; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line)

=>

IT

```
116:210833
DN
     Structures of cytotoxic substances and new quinazoline derivatives
ΤI
     produced by a fungus from a saltwater fish
     Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika;
     Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi,
     Hirofumi; Shingu, Tetsuro
CS
     Osaka Univ. Pharm. Sci., Osaka, Japan
     Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30
SO
     CODEN: TYKYDS
DT
     Journal
LΑ
     Japanese
```

10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1

CC

GI

AB Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior.

ST Aspergillus fumiquinazoline cytotoxicity structure

IT Nomenclature, new natural products

(fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Aspergillus fumigatus

(fumiquinazolines from, structure and cytotoxicity of)

IT Molecular structure, natural product

(of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline E (quinazoline), from Aspergillus fumigatus)

ΙT Neoplasm inhibitors (leukemia, fumiquinazolines as, from Aspergillus fumigatus) ΙT 140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) 67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. IT 12771-72-1, Verruculogen 51177-07-2 62867-47-4, Fumigaclavine C 74149-38-5 111427-99-7, TR 3 111468-06-5 115589-18-9 118974-02-0, Fumitremorgin 137494-04-3 140715-85-1, Fumiquinazoline A 140715-86-2, Fumiquinazoline D 140715-87-3, Fumiquinazoline E Fumiquinazoline B 140924-01-2, Fumiquinazoline C RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (structure and cytotoxic activity of, from Aspergillus fumigatus)

=>

```
DN
     124:285652
ΤI
     Alterations in expression of the multidrug resistance-associated
     protein (MRP) gene in high-grade transitional cell carcinoma of
     the bladder
     Clifford, S. C.; Neal, D. E.; Lunec, J.
ΑU
CS
     Medical School, University of Newcastle-upon-Tyne, Newcastle-upon-Tyne,
     NE2 4HH, UK
     British Journal of Cancer (1996), 73(5), 659-66
SO
     CODEN: BJCAAI; ISSN: 0007-0920
PB
     Stockton
DT
     Journal
LΑ
     English
CC
     14-1 (Mammalian Pathological Biochemistry)
     Section cross-reference(s): 1
     Expression of the MRP gene has been demonstrated in vitro to be
AB
     a causal factor in non-P-glycoprotein-mediated multidrug
     resistance, and is implicated in resistance to a no. of
     the chemotherapeutic agents currently used in the treatment of high-grade
     transitional cell carcinoma (TCC) of the bladder (doxorubicin, epirubicin
     and vinblastine). Using a sensitive RT-PCR-based technique, we have
     quantified MRP mRNA levels in a series of untreated TCC (n=24),
     normal bladder (n=5) and control tissue and cell line samples.
     MRP mRNA was widely expressed and detectable in all samples
     analyzed, with considerable (up to 190-fold) variation obsd. between
     individual tumor samples. MRP mRNA levels found in TCC samples
     were lower than those detd. for normal peripheral mononucleocyte
     (2.3-fold) and testis (4.1-fold) samples, previously reported to be
     high-expressing tissues, and varied over a similar range to that obsd. in normal bladder samples. Results indicate that MRP mRNA levels
     in a greater proportion of high-grade (G3) bladder tumors (55%, 6/11) are
     significantly reduced (P=0.018) compared with low- and moderate-grade
     (G1/2) bladder tumors (8%, 1/13), and suggest that MRP mRNA
     levels frequently become reduced as a consequence of tumor progression to
     advanced, poorly differentiated disease. No correlation was apparent
     between MRP and MDR1 mRNA levels, thus providing no evidence to
     suggest common regulation of the two genes. In a limited no. of patients,
     no evidence was found to support a role for MRP mRNA levels as a
     determinant of response to chemotherapy in patients being uniformly
     treated with either cisplatin-methotrexate-vinblastine (n=6) or
     epirubicin-cisplatin-methotrexate (n=4) regimens. Similarly, no overall
     pattern of altered MRP mRNA expression was obsd. following
     chemotherapy in four patients from whom post chemotherapy biopsies were
     taken. This study provides a useful pilot investigation regarding the
     level, variation and pattern of MRP mRNA expression in TCC of
     the bladder, and suggests that further studies to establish the clin.
     significance of these variations are required.
ST
     multidrug resistance protein gene bladder carcinoma
IT
     Gene, animal
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (MRP; multidrug resistance-assocd. protein gene
        expression in human high-grade transitional cell carcinoma of the
        bladder)
IT
     Proteins, specific or class
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (multidrug resistance-assocd. protein (MRP);
        multidrug resistance-assocd. protein gene expression in human
        high-grade transitional cell carcinoma of the bladder)
IT
     Neoplasm inhibitors
        (multidrug resistance-assocd. protein gene expression in
        human high-grade transitional cell carcinoma of the bladder)
IT
     Ribonucleic acids, messenger
```

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

IT Bladder

=>

(neoplasm, transitional cell carcinoma, multidrug resistance -assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

IT 59-05-2, Methotrexate 865-21-4, Vinblastine **15663-27-1**, Cisplatin 56420-45-2, Epirubicin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(multidrug resistance-assocd. protein gene expression in human high-grade transitional cell carcinoma of the bladder)

116:210833 DN ΤI

Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish

Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; ΑU Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro

Osaka Univ. Pharm. Sci., Osaka, Japan CS

SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS

DT Journal

LA Japanese

CC 10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1

GΙ

AΒ Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior.

ST Aspergillus fumiquinazoline cytotoxicity structure

ΙT Nomenclature, new natural products

(fumiquinazoline A (quinazoline), from Aspergillus fumiqatus)

ΙT Aspergillus fumigatus

(fumiquinazolines from, structure and cytotoxicity of)

ΙT Molecular structure, natural product

(of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

ΙT Molecular structure, natural product

(of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline E (quinazoline), from Aspergillus fumigatus)

ΙT Neoplasm inhibitors (leukemia, fumiquinazolines as, from Aspergillus fumigatus) ΙT 140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) IT67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. 12771-72-1, Verruculogen 51177-07-2 62867-47-4, Fumigaclavine C 74149-38-5 111427-99-7, TR 3 111468-06-5 115589-18-9 118974-02-0, Fumitremorgin 137494-04-3 140715-85-1, Fumiguinazoline A 140715-86-2, Fumiquinazoline D 140715-87-3, Fumiquinazoline E Fumiquinazoline B 140924-01-2, Fumiquinazoline C 140852-71-7, RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (structure and cytotoxic activity of, from Aspergillus fumigatus)

=>

AN 1994:94935 CAPLUS

DN 120:94935

TI Cross-resistance to diverse drugs is associated with primary cisplatin resistance in ovarian cancer cell lines

AU Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter J.; Ozols, Robert F.; Hamilton, Thomas C.

CS Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA

SO Cancer Research (1993), 53(21), 5225-32 CODEN: CNREA8; ISSN: 0008-5472

DT Journal

LA English

CC 1-6 (Pharmacology)

AB The authors have previously obtained, by exposure to near continuous increasing concns. of cisplatin, a panel of human ovarian cancer cell lines that exhibit a wide range of primary resistance to the drug (9- to >400-fold). These cells had strikingly increased (4- to 50-fold) levels of glutathione (GSH) as compared with the drug-sensitive cells of origin (A. K. Godwin et al., Proc. Natl. Acad. Sci. USA, 89: 3070-3074, 1992). Using this panel of resistant cell lines, the authors evaluated cross-resistance to classical alkylating agents, natural product drugs, and irradn. Cross-resistance to carboplatin paralleled that of cisplatin, culminating is .apprx. 250-fold resistance. Similarly, melphalan cross-resistance continued to increase to >400-fold and again paralleled the primary cisplatin resistance. Cell lines with low to very high levels of resistance to cisplatin are 8-850-fold resistant to the epipodophyllotoxin deriv. etoposide. Cross-resistance is also obsd. for other natural product drugs, including Adriamycin (.apprx.80-fold), mitoxantrone (.apprx.440-fold), and taxol (.apprx.40-fold). Cross-resistance to irradn. is, however, modest (<2-fold). The cells with greatest primary resistance to cisplatin most commonly had the highest cross-resistance to the other drugs examd. The cross-resistance to the natural produce category drugs was found not to be mediated by the products of either the multidrug resistance 1 (MDR1) or multidrug resistance-assocd. protein (MRP) genes based on lack of coordinate increased expression or amplification of these genes as assessed by Northern and Southern blot analyses. Also, verapamil failed to markedly increase drug sensitivity. Although there was no indication that these natural product drug efflux pumps were operative, the authors obsd. decreased doxorubicin accumulation in these cell lines cross-resistant to natural products. Alternations in DNA topoisomerase II mRNA levels, which were obsd. in human tumor cell lines selected in vitro for resistance to etoposide or teniposide, were not detected. Only intracellular levels of GSH correlated with cross-resistance to these diverse anticancer agents and partial loss of resistance was assocd. with a marked decrease in glutathione levels. In the absence of alternative mechanisms, the authors speculate that the very broad clin. relevant cross-resistance seen in this model system may, at least in part, be the direct result of GSH-mediated drug inactivation or may be due to a combination of GSH conjugation to drug and conjugate efflux mediated by the putative ATP-dependent glutathione S-conjugate export pump.

ST cisplatin resistance neoplasm cross resistance glutathione

IT Neoplasm inhibitors

(cisplatin as, resistance to, cross-resistance to, in humans cells, GSH in mechanism of)

IT Radiation

(cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of)

IT Biological transport

(of doxorubicin, in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in)

IT Drug resistance

(to cisplatin, cross-resistance to, in neoplasm of humans, GSH in

mechanism of) IT 148-82-3, Melphalan 33069-62-4, Taxol 33419-42-0, VP-16 41575-94-4, Carboplatin 65271-80-9, Mitoxantrone RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of) ΙT 23214-92-8, Adriamycin RL: PRP (Properties) (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of, transport in relation to) ΙT 70-18-8, Glutathione, biological studies RL: BIOL (Biological study) (in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in) IT 15663-27-1, Cisplatin RL: BIOL (Biological study) (resistance to, in neoplasm of humans, cross-resistance in, GSH in

=>

mechanism of)

```
1996:646439 CAPLUS
ΑN
DN
     125:266006
     Use of protein kinase inhibitors in preventing multidrug
TI
     resistance in cancer cells
IN
     Chaudhary, Preet; Shtil, Alexander A.; Roninson, Igor B.
PA
     Board of Trustees of the University of Illinois, USA
SO
     PCT Int. Appl., 75 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
IC
     ICM A61K045-06
CC
     1-6 (Pharmacology)
FAN.CNT 4
     PATENT NO.
                      KIND DATE
                                          APPLICATION NO.
                                                            DATE
                                           -----
PΙ
     WO 9625949
                       A1
                            19960829
                                           WO 1996-US422
                                                            19960111
         W: CA, JP
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     US 5972598
                           19991026
                                           US 1995-370724
                                                            19950110
                       A
     EP 804240
                                           EP 1996-903458
                      A1
                           19971105
                                                            19960111
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE
     JP 10512277
                      Т2
                                          JP 1996-522483
                           19981124
                                                            19960111
     US 6171786
                                           US 1996-659877
                      В1
                            20010109
                                                            19960607
PRAI US 1995-370724
                      Α
                            19950110
     US 1992-947659
                      В2
                            19920918
     WO 1996-US422
                      W
                            19960111
     Methods are disclosed for preventing the emergence of multidrug
AΒ
     resistance in tumor cells during cancer chemotherapy. In
     particular, protein kinase inhibitors are used to prevent the induction of
     expression of the multidrug resistance gene (MDR1) encoding
     P-glycoprotein by chemotherapeutic drugs. MDR1 expression, which results
     in tumor cell resistance to subsequent treatment with certain
     chemotherapeutic drugs, is shown herein to be induced in response to
     treatment with various cytotoxic agents, including such agents that are
     and are not substrates for P-glycoprotein-mediated efflux from cancer
     cells. Inhibitors of protein kinases, in particular protein kinase C, are
     shown to suppress this cellular response. In addn., such protein kinase
     inhibitors are also shown to inhibit expression of a gene encoding a
     multidrug resistance-assocd. protein (the MRP gene).
     Methods are disclosed for using such protein kinase inhibitors to both
     suppress induction of MDR1 gene expression by cytotoxic drugs and to
     inhibit expression of MRP. Also provided are methods for
     identifying protein kinase inhibitors that have either or both of these
     effects on MDR1 and MRP expression. Thus, the invention
     provides useful methods and reagents for preventing the emergence of
     multidrug resistance in tumor cells treated with cytotoxic and
     chemotherapeutic drugs in cancer patients undergoing chemotherapy, when
     such protein kinase inhibitors are administered prior to or simultaneous
     with cytotoxic drug treatment in such individuals.
     protein kinase inhibitor multidrug resistance inhibition; MDR
ST
     inhibition protein kinase inhibitor; cancer therapy protein kinase
     inhibitor MDR
IT
     Gene, animal
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (MRP; protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells) .
IT
    Lymphocyte
        (differentiation; protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
    Cell differentiation
        (lymphoid cell; protein kinase inhibitors for prevention of multidrug
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resistance in cancer cells)
ΙT
     Proteins, specific or class, biological studies
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (multidrug resistance-assocd.; protein kinase inhibitors for
        prevention of multidrug resistance in cancer cells)
IT
     Biological transport
     Cytotoxic agents
     HeLa cell
     Lymphoma
     Neoplasm inhibitors
        (protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Glycophosphoproteins .
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (P-, gene mdrl, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
        (T-cell, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
        (acute monocytic, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
        (acute myelogenous, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
     Uterus, neoplasm
        (cervix, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Therapeutics
        (chemo-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
        (chronic myelocytic, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
     Skin, neoplasm
        (epidermoid carcinoma, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
        (fibro-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Neoplasm inhibitors
        (hematol., protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Neoplasm inhibitors
        (leukemia, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
    Gene, animal
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (mdrl, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
ΙT
     Drug resistance
        (multi-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Bladder
     Mammary gland
        (neoplasm, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
ΙT
     Thymus gland
        (neoplasm, thymoma, protein kinase inhibitors for prevention of
```

multidrug resistance in cancer cells)

IT Leukemia

(promyelocytic, protein kinase inhibitors for prevention of multidrug resistance in cancer cells)

IT Neoplasm inhibitors

(solid, protein kinase inhibitors for prevention of multidrug resistance in cancer cells)

IT Light

=>

(white, and calphostin C; protein kinase inhibitors for prevention of multidrug resistance in cancer cells)

59-05-2, Methotrexate 127-07-1, Hydroxyurea 147-94-4, Cytosine IT arabinoside 305-03-3, Chlorambucil 446-72-0, Genistein 865-21-4, Vinblastine 1405-10-3, Neomycin sulfate 1405-10-3D, Neomycin sulfate, derivs. 15663-27-1, Cisplatin 20830-81-3, Daunorubicin 25316-40-9, Adriamycin 34316-15-9, Chelerythrine 34316-15-9D, Chelerythrine, derivs. 62996-74-1, Staurosporine 62996-74-1D, 63177-57-1, Methyl 2,5-dihydroxycinnamate Staurosporine, derivs. 84477-87-2, H7 84477-87-2D, H7, derivs. 70563-58-5, Herbimycin A 91742-10-8, HA1004 100827-28-9, Erbstatin 88494-43-3 100827-28-9D, Erbstatin, derivs. 118409-58-8, Tyrphostin A25 118409-58-8D, Tyrphostin A25, derivs. 121263-19-2, Calphostin C 121263-19-2D, Calphostin C, derivs. 149092-34-2, Tyrphostin B46 149092-34-2D, Tyrphostin B46, derivs.

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(protein kinase inhibitors for prevention of multidrug resistance in cancer cells)

IT 9026-43-1, Protein kinase 141436-78-4, Protein kinase C
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
 (Biological study); PROC (Process)

(protein kinase inhibitors for prevention of multidrug resistance in cancer cells)



L5 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2003 ACS

AN 1992:210833 CAPLUS

DN 116:210833

TI Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish

AU Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro

CS Osaka Univ. Pharm. Sci., Osaka, Japan

SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS

DT Journal

LA Japanese

CC 10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1

GI

AB Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior.

ST Aspergillus fumiquinazoline cytotoxicity structure

IT Nomenclature, new natural products

(fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Aspergillus fumigatus

(fumiquinazolines from, structure and cytotoxicity of)

IT Molecular structure, natural product

(of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)

· IT Molecular structure, natural product

(of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline E (quinazoline), from Aspergillus fumigatus) ITNeoplasm inhibitors (leukemia, fumiquinazolines as, from Aspergillus fumigatus) IT140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) IT 67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. 12771-72-1,. Verruculogen 51177-07-2 62867-47-4, Fumigaclavine C 74149-38-5 **111427-99-7**, TR 3 111468-06-5 115589-18-9 118974-02-0 , Fumitremorgin C 137494-04-3 140715-85-1, Fumiquinazoline A 140715-86-2, Fumiquinazoline D 140852-71-7, Fumiquinazoline B 140715-87-3, Fumiquinazoline E 140924-01-2, Fumiquinazoline C RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (structure and cytotoxic activity of, from Aspergillus fumigatus)

AN 1994:94935 CAPLUS

DN 120:94935

TI Cross-resistance to diverse drugs is associated with primary cisplatin resistance in ovarian cancer cell lines

AU Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter J.; Ozols, Robert F.; Hamilton, Thomas C.

CS Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA

SO Cancer Research (1993), 53(21), 5225-32 CODEN: CNREA8; ISSN: 0008-5472

DT Journal

LA English

CC 1-6 (Pharmacology)

AB The authors have previously obtained, by exposure to near continuous increasing concns. of cisplatin, a panel of human ovarian cancer cell lines that exhibit a wide range of primary resistance to the drug (9- to >400-fold). These cells had strikingly increased (4- to 50-fold) levels of glutathione (GSH) as compared with the drug-sensitive cells of origin (A. K. Godwin et al., Proc. Natl. Acad. Sci. USA, 89: 3070-3074, 1992). Using this panel of resistant cell lines, the authors evaluated cross-resistance to classical alkylating agents, natural product drugs, and irradn. Cross-resistance to carboplatin paralleled that of cisplatin, culminating is .apprx. 250-fold resistance. Similarly, melphalan cross-resistance continued to increase to >400-fold and again paralleled the primary cisplatin resistance. Cell lines with low to very high levels of resistance to cisplatin are 8-850-fold resistant to the epipodophyllotoxin deriv. etoposide. Cross-resistance is also obsd. for other natural product drugs, including Adriamycin (.apprx.80-fold), mitoxantrone (.apprx.440-fold), and taxol (.apprx.40-fold). Cross-resistance to irradn. is, however, modest (<2-fold). The cells with greatest primary resistance to cisplatin most commonly had the highest cross-resistance to the other drugs examd. The cross-resistance to the natural produce category drugs was found not to be mediated by the products of either the multidrug resistance 1 (MDR1) or multidrug resistance-assocd. protein (MRP) genes based on lack of coordinate increased expression or amplification of these genes as assessed by Northern and Southern blot analyses. Also, verapamil failed to markedly increase drug sensitivity. Although there was no indication that these natural product drug efflux pumps were operative, the authors obsd. decreased doxorubicin accumulation in these cell lines cross-resistant to natural products. Alternations in DNA topoisomerase II mRNA levels, which were obsd. in human tumor cell lines selected in vitro for resistance to etoposide or teniposide, were not detected. Only intracellular levels of GSH correlated with cross-resistance to these diverse anticancer agents and partial loss of resistance was assocd. with a marked decrease in glutathione levels. In the absence of alternative mechanisms, the authors speculate that the very broad clin. relevant cross-resistance seen in this model system may, at least in part, be the direct result of GSH-mediated drug inactivation or may be due to a combination of GSH conjugation to drug and conjugate efflux mediated by the putative ATP-dependent glutathione S-conjugate export pump.

ST cisplatin resistance neoplasm cross resistance glutathione

IT Neoplasm inhibitors

(cisplatin as, resistance to, cross-resistance to, in humans cells, GSH in mechanism of)

IT Radiation

(cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of)

IT Biological transport

(of doxorubicin, in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in)

IT Drug resistance

(to cisplatin, cross-resistance to, in neoplasm of humans, GSH in

mechanism of)

IT 148-82-3, Melphalan 33069-62-4, Taxol 33419-42-0, VP-16 41575-94-4, Carboplatin 65271-80-9, Mitoxantrone

RL: PRP (Properties)
 (cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of)

IT 23214-92-8, Adriamycin RL: PRP (Properties)

(cross-resistance of, in neoplasm of humans, to cisplatin resistance, GSH in mechanism of, transport in relation to)

IT 70-18-8, Glutathione, biological studies

RL: BIOL (Biological study)

(in neoplasm of humans resistance to cisplatin, cross-resistance to other agents in)

IT 15663-27-1, Cisplatin

RL: BIOL (Biological study)

(resistance to, in neoplasm of humans, cross-resistance in, GSH in mechanism of)

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CY SUBSCRIBER PRICE

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STN INTERNATIONAL LOGOFF AT 16:14:51 ON 19 JUN 2003

L10 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2003 ACS

AN 1992:210833 CAPLUS

DN 116:210833

TI Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish

AU Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro

CS Osaka Univ. Pharm. Sci., Osaka, Japan

SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS

DT Journal

LA Japanese

CC 10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1

GI

AB Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior.

ST Aspergillus fumiquinazoline cytotoxicity structure

IT Nomenclature, new natural products

(fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Aspergillus' fumigatus

(fumiquinazolines from, structure and cytotoxicity of)

IT Molecular structure, natural product

(of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline E (quinazoline), from Aspergillus fumigatus)

IT' Neoplasm inhibitors (leukemia, fumiquinazolines as, from Aspergillus fumigatus) IT140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) IT 67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. 12771-72-1, Verruculogen 51177-07-2 62867-47-4, Fumigaclavine C 74149-38-5 115589-18-9 **118974-02-0** 140715-85-1, Fumiquinazoline A **111427-99-7**, TR 3 111468-06-5 , Fumitremorgin C 137494-04-3 140715-86-2, Fumiquinazoline D 140852-71-7, Fumiquinazoline B 140715-87-3, Fumiquinazoline E 140924-01-2, Fumiquinazoline C RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (structure and cytotoxic activity of, from Aspergillus fumigatus)

=>

=> d 112 129 all

- L12 ANSWER 129 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1994:94935 CAPLUS
- DN 120:94935
- TI Cross-resistance to diverse drugs is associated with primary cisplatin resistance in ovarian cancer cell lines
- AU Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter J.; Ozols, Robert F.; Hamilton, Thomas C.
- CS Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA
- SO Cancer Research (1993), 53(21), 5225-32 CODEN: CNREA8; ISSN: 0008-5472
- DT Journal
- LA English
- CC 1-6 (Pharmacology)
- AΒ The authors have previously obtained, by exposure to near continuous increasing concns. of cisplatin, a panel of human ovarian cancer cell lines that exhibit a wide range of primary resistance to the drug (9- to >400-fold). These cells had strikingly increased (4- to 50-fold) levels of glutathione (GSH) as compared with the drug-sensitive cells of origin (A. K. Godwin et al., Proc. Natl. Acad. Sci. USA, 89: 3070-3074, 1992). Using this panel of resistant cell lines, the authors evaluated cross-resistance to classical alkylating agents, natural product drugs, and irradn. Cross-resistance to carboplatin paralleled that of cisplatin, culminating is .apprx. 250-fold resistance. Similarly, melphalan cross-resistance continued to increase to >400-fold and again paralleled the primary cisplatin resistance. Cell lines with low to very high levels of resistance to cisplatin are 8-850-fold resistant to the epipodophyllotoxin deriv. etoposide. Cross-resistance is also obsd. for other natural product drugs, including Adriamycin (.apprx.80-fold), mitoxantrone (.apprx.440-fold), and taxol (.apprx.40-fold). Cross-resistance to irradn. is, however, modest (<2-fold). The cells with greatest primary resistance to cisplatin most commonly had the highest cross-resistance to the other drugs examd. The cross-resistance to the natural produce category drugs was found not to be mediated by the products of either the multidrug resistance 1 (MDR1) or multidrug resistance -assocd. protein (MRP) genes based on lack of coordinate increased expression or amplification of these genes as assessed by Northern and Southern blot analyses. Also, verapamil failed to markedly increase drug sensitivity. Although there was no indication that these natural product drug efflux pumps were operative, the authors obsd. decreased doxorubicin accumulation in these cell lines cross-resistant to natural products. Alternations in DNA topoisomerase II mRNA levels, which were obsd. in human tumor cell lines selected in vitro for resistance to etoposide or teniposide, were not detected. Only intracellular levels of GSH correlated with cross-resistance to these diverse anticancer agents and partial loss of resistance was assocd. with a marked decrease in glutathione levels. In the absence of alternative mechanisms, the authors speculate that the very broad clin. relevant cross-resistance seen in this model system may, at least in part, be the direct result of GSH-mediated drug inactivation or may be due to a combination of GSH conjugation to drug and conjugate efflux mediated by the putative ATP-dependent glutathione S-conjugate export pump.
- ST cisplatin resistance neoplasm cross resistance glutathione
- IT Neoplasm inhibitors

```
(cisplatin as, resistance to, cross-resistance to,
         in humans cells, GSH in mechanism of)
IT
     Radiation
         (cross-resistance of, in neoplasm of humans, to cisplatin
         resistance, GSH in mechanism of)
IT
     Biological transport
         (of doxorubicin, in neoplasm of humans resistance to
         cisplatin, cross-resistance to other agents in)
IT
     Drug resistance
         (to cisplatin, cross-resistance to, in neoplasm of humans,
         GSH in mechanism of)
IT
     148-82-3, Melphalan
                            33069-62-4, Taxol
                                                33419-42-0, VP-16
                                                                     41575-94-4,
     Carboplatin
                   65271-80-9, Mitoxantrone
     RL: PRP (Properties)
         (cross-resistance of, in neoplasm of humans, to cisplatin
        resistance, GSH in mechanism of)
IT
     23214-92-8, Adriamycin
     RL: PRP (Properties)
         (cross-resistance of, in neoplasm of humans, to cisplatin
        resistance, GSH in mechanism of, transport in relation to)
IT
     70-18-8, Glutathione, biological studies
     RL: BIOL (Biological study)
         (in neoplasm of humans resistance to cisplatin, cross-
        resistance to other agents in)
IT
     15663-27-1, Cisplatin
     RL: BIOL (Biological study)
         (resistance to, in neoplasm of humans, cross-
        resistance in, GSH in mechanism of)
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L3
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L4
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          14128 S L2
L6
             53 S L3
L7
              2 S L5 AND L6
                E RESISTANCE
L8
         901098 S E3
             15 S L8 AND L6
L9
           2634 S L5 AND L8
L10
L11
           2146 S MRP
L12
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Executing the logoff script...

CC 1-6 (Pharmacology) ΑB A human bladder cancer cell line resistant to adriamycin (ADM), T24/ADM9 has been established in vitro by exposing T24 parent cells to progressively higher concns. of the drug over a period of 12 mo. T24/ADM9 cells were 9 times more resistant to ADM than the T24 parent, and showed various degrees of cross-resistance to an ADM deriv., vinca alkaloids and a DNA topoisomerase II (Topo II)-targeting agent, etoposide. No significant difference was obsd. in the cellular accumulation of ADM between the T24/ADM9 and T24 parent cells. A Northern blot anal. showed an overexpression of multidrug resistance -assocd. protein (MRP) mRNA, but no overexpression of multidrug ${f resistance}^{-1}$ (MDR1) mRNA was obsd. in the T24/ADM9 cells. A flow cytometric anal. showed that the MDR1 gene product, P-glycoprotein (Pgp), is not expressed on the T24/ADM9 cells. T24/ADM9 showed approx. the parental level of DNA Topo II catalytic activity. In Western blot and Northern blot analyses, however, the cellular level of DNA Topo II was apparently much lower in T24/ADM9 than in the T24 parent. Thus, these results suggest that a decreased cellular level of DNA Topo II and an overexpression of MRP gene may be responsible for the expression of an MDR phenotype in the T24/ADM9 cells and that such non-Pgp-mediated, atypical MDR may develop in bladder cancer treated with chemotherapy including ADM. STatypical multidrug resistant bladder cancer cell ITProteins, specific or class RL: BSU (Biological study, unclassified); BIOL (Biological study) (multidrug resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Gene, animal RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Neoplasm inhibitors (bladder carcinoma, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT Drug resistance (multi-, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT Bladder (neoplasm, carcinoma, inhibitors, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) 50-07-7, Mitomycin C 51-21-8, 5 Fluorouracil 57-22-7, Vincristine ΙT 865-21-4, Vinblastine **15663-27-1**, Cisplatin 33419-42-0, Etoposide 56420-45-2, Epirubicin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cross-resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT142805-56-9, DNA topoisomerase II RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to decreased level of DNA Topo II) IT 25316-40-9, Adriamycin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (resistance to; non-P-glycoprotein-mediated atypical

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     JP 2002502592
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                            19980205
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                            19990205
     WO 1999-US2577
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              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 3
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 14 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     1999:173577 CAPLUS
DN
     131:13477
     Multiple mechanisms confer drug resistance to mitoxantrone in
TΙ
     the human 8226 myeloma cell line
     Hazlehurst, Lori A.; Foley, Nils E.; Gleason-Guzman, Mary C.; Hacker,
AU
     Miles P.; Cress, Anne E.; Greenberger, Lee W.; De Jong, Mariska C.;
    Dalton, William S.
     Department of Biochemistry, Pharmacology, and Internal Medicine, H. Lee
CS
     Moffitt Cancer Center, University of South Florida, Tampa, FL, 33612, USA
SO
     Cancer Research (1999), 59(5), 1021-1028
     CODEN: CNREA8; ISSN: 0008-5472
PB
     AACR Subscription Office
DT
     Journal
LΑ
     English
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              ALL CITATIONS AVAILABLE IN THE RE FORMAT
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     ANSWER 15 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     1999:2922 CAPLUS
DN
     130:177236
     Reversal of a novel multidrug resistance mechanism in human
     colon carcinoma cells by fumitremorgin C
     Rabindran, Sridhar K.; He, Haiyin; Singh, Maya; Brown, Eileen; Collins,
ΑU
     Karen I.; Annable, Tami; Greenberger, Lee M.
CS
     Oncology and Immunology Research, Wyeth-Ayerst Research, Pearl River, NY,
     10965, USA
SO
     Cancer Research (1998), 58(24), 5850-5858
     CODEN: CNREA8; ISSN: 0008-5472
     AACR Subscription Office
PB
DT
     Journal
     English
              THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD
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     FILE 'REGISTRY' ENTERED AT 15:55:33 ON 19 JUN 2003
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L3
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L4
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L7
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                E RESISTANCE
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L8
L9
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=> s 15 and 18
L10
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=> s mrp
L11
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=> s 110 and 111
           129 L10 AND L11
=> d 112 100-129
L12 ANSWER 100 OF 129 CAPLUS COPYRIGHT 2003 ACS
     1997:561610 CAPLUS
DN
     127:214731
ΤI
     Molecular targeting of mitomycin C chemotherapy
     Nishiyama, Masahiko; Suzuki, Katsuyuki; Kumazaki, Tsutomu; Yamamoto,
     Wataru; Toge, Tetsuya; Okamura, Tatsunori; Kurisu, Kaoru
     Department of Biochemistry and Biophysics, Research Institute for
CS
     Radiation Biology and Medicine, Hiroshima University, Hiroshima, 734,
     Japan
SO
     International Journal of Cancer (1997), 72(4), 649-656
     CODEN: IJCNAW; ISSN: 0020-7136
PB
     Wilev-Liss
DT
     Journal
LA
     English
L12
     ANSWER 101 OF 129 CAPLUS COPYRIGHT 2003 ACS
     1997:560303 CAPLUS
AN
DN
     127:242890
ΤI
     Analysis of expression of cMOAT (MRP2), MRP3, MRP4, and MRP5, homologs of
     the multidrug resistance-associated protein gene (MRP1), in
     human cancer cell lines
ΑU
     Kool, Marcel; De Haas, Marcel; Scheffer, George L.; Scheper, Rik J.; Van
     Eijk, Michiel J. T.; Juijn, Jenneke A.; Baas, Frank; Borst, Piet
CS
     Division of Molecular Biology, The Netherlands Cancer Institute,
     Amsterdam, 1066 CX, Neth.
SO
     Cancer Research (1997), 57(16), 3537-3547
     CODEN: CNREA8; ISSN: 0008-5472
PB
     American Association for Cancer Research
DT
     Journal
LΑ
     English
     ANSWER 102 OF 129 CAPLUS COPYRIGHT 2003 ACS
L12
     1997:548817 CAPLUS
AN
DN
     127:214767
     A novel quinoline derivative, MS-209, overcomes drug resistance
TΙ
     of human lung cancer dells expressing the multidrug resistance
     -associated protein (MRP) gene
ΑU
     Narasaki, Fumihiko; Oka, Mikio; Fukuda, Minoru; Nakano, Reiji; Ikeda,
     Koki; Takatani, Hiroshi; Terashi, Kenji; Soda, Hiroshi; Yano, Osamu;
     Nakamura, Tsuyoshi; Doyle, L. Austin; Tsuruo, Takashi; Kohno, Shigeru
     School Medicine, Nagasaki University, Nagasaki, 852, Japan
CS
SO
     Cancer Chemotherapy and Pharmacology (1997), 40(5), 425-432
     CODEN: CCPHDZ; ISSN: 0344-5704
PΒ
     Springer
DT
     Journal
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LΑ

English

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L12 ANSWER 103 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1997:432371 CAPLUS
DN
     127:104013
     Isolation from a human MDR lung cell line of multiple clonal
TI
     subpopulations which exhibit significantly different drug
     Heenan, Mary; O'driscoll, Lorraine; Cleary, Irene; Connolly, Lisa; Clynes,
ΑU
     Martin
CS
     National Cell and Tissue Culture Centre/BioResearch Ireland, Dublin City
     University, Dublin, 9, Ire.
SO
     International Journal of Cancer (1997), 71(5), 907-915
     CODEN: IJCNAW; ISSN: 0020-7136
PB
     Wiley-Liss
     Journal
DT
LA
     English
L12 ANSWER 104 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1997:343754 CAPLUS
     127:44550
DN
     Lack of a point mutation of human DNA topoisomerase II in
ΤI
     multidrug-resistant anaplastic thyroid carcinoma cell lines
ΑU
     Satake, Shoji; Sugawara, Isamu; Watanabe, Masatoshi; Takami, Hiroshi
     Dep. Surgery, Teikyo Univ. Sch. Med., Tokyo, 171, Japan
CS
     Cancer Letters (Shannon, Ireland) (1997), 116(1), 33-39
SO
     CODEN: CALEDQ; ISSN: 0304-3835
PB
     Elsevier
DT
     Journal
LΆ
     English
     ANSWER 105 OF 129 CAPLUS COPYRIGHT 2003 ACS
L12
     1997:316426 CAPLUS
ΑN
DN
     126:338501
TI
     Intermittent exposure to doxorubicin in vitro selects for multifactorial
     non-P-glycoprotein-associated multidrug resistance in RPMI 8226
     human myeloma cells
ΑU
     Wyler, Beat; Shao, Ying; Schneider, Erasmus; Cianfriglia, Maurizio;
     Scheper, Rik J.; Frey, Beat M.; Gieseler, Frank; Schmid, Luzius;
     Twentyman, Peter R.; Lehnert, Manfred
CS
     Cancer Research Laboratory, Department C of Internal Medicine,
     Kantonsspital St Gallen, St Gallen, 9007, Switz.
SO
     British Journal of Haematology (1997), 97(1), 65-75
     CODEN: BJHEAL; ISSN: 0007-1048
PB
     Blackwell
DT
     Journal .
LΑ
     English
L12
    ANSWER 106 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1997:266139 CAPLUS
DN
     126:311883
     Possible role of the multidrug resistance-associated protein (
ΤI
     MRP) in chemoresistance of human melanoma cells
     Berger, Walter; Hauptmann, Erich; Elbling, Leonilla; Vetterlein, Monika;
ΑU
     Kokoschka, Eva M.; Micksche, Michael
CS
     Dep. Applied & Experimental Oncology, Inst. Tumor Biology/Cancer Res.,
     Vienna Univ., Vienna, Austria
SO
     International Journal of Cancer (1997), 71(1), 108-115
     CODEN: IJCNAW; ISSN: 0020-7136
PB
     Wiley-Liss
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DT

LΑ

Journal

English

- L12 ANSWER 107 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1997:242551 CAPLUS
- DN 126:311861
- TI Carbamoylation of glutathione reductase by N,N-bis(2-chloroethyl)-N-nitrosourea associated with inhibition of multidrug resistance protein (MRP) function
- AU Vanhoefer, Udo; Yin, Ming-Biao; Harstrick, Andreas; Seeber, Siegfried; Rustum, Youcef M.
- CS DEPARTMENT OF EXPERIMENTAL THERAPEUTICS, GRACE CANCER DRUG CENTER, ROSWELL PARK CANCER INSTITUTE, BUFFALO, NY, 14263, USA
- SO Biochemical Pharmacology (1997), 53(6), 801-809 CODEN: BCPCA6; ISSN: 0006-2952
- PB Elsevier
- DT Journal
- LA English
- L12 ANSWER 108 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1997:233235 CAPLUS
- DN 126:271951
- TI In vitro cross-resistance and collateral sensitivity in seven resistant small-cell lung cancer cell lines: preclinical identification of suitable drug partners to taxotere, taxol, topotecan and gemcitabine
- AU Jensen, P. B.; Holm, B.; Sorensen, M.; Christensen, I. J.; Sehested, M.
- CS Laboratory of Experimental Medical Oncology, The Finsen Center, Rigshospitalet, Copenhagen, DK-2100, Den.
- SO British Journal of Cancer (1997), 75(6), 869-877 CODEN: BJCAAI; ISSN: 0007-0920
- PB Churchill Livingstone
- DT Journal
- LA English
- L12 ANSWER 109 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1997:154474 CAPLUS
- DN 126:207244
- In vitro evaluation of new anticancer drugs, exemplified by vinorelbine, using the fluorometric microculture cytotoxicity assay on human tumor cell lines and patient biopsy cells
- AU Fridborg, Helena; Nygren, Peter; Dhar, Sumeer; Csoka, Katalin; Kristensen, Joergen; Larsson, Rolf
- CS Division of Clinical Pharmacology, University Hospital, Uppsala University, Uppsala, S-751 85, Swed.
- SO Journal of Experimental Therapeutics & Oncology (1996), 1(5), 286-295 CODEN: JETOFX; ISSN: 1359-4117
- PB Rapid Science Publishers
- DT Journal
- LA English
- L12 ANSWER 110 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1997:92153 CAPLUS
- DN 126:112882
- TI Rapid recovery of a functional MDR phenotype caused by **MRP** after a transient exposure to MDR drugs in a revertant human lung cancer cell line
- AU Manzano, R. Gonzalez; Versanvoort, C.; Wright, K.; Twentyman, P.R.
- CS Clinical Oncology and Radiotherapeutics Unit Medical Research Council Centre, Cambridge, 2QH, UK
- SO European Journal of Cancer, Part A (1996), 32A(12), 2136-2141 CODEN: EJCTEA
- PB Elsevier
- DT Journal
- LA English

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L12
     ANSWER 111 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1997:24455 CAPLUS
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     126:112834
     The multidrug resistance-associated protein gene confers drug
TI
     resistance in human gastric and colon cancers
     Tomonaga, Michio; Oka, Mikio; Narasaki, Fumihiko; Fukuda, Minoru; Nakano,
     Reiji; Takatani, Hiroshi; Ikeda, Koki; Terashi, Kenji; Matsuo, Isao; et
CS
     The Second Department of Internal Medicine, Nagasaki University School of
     Medicine, Nagasaki, 852, Japan
     Japanese Journal of Cancer Research (1996), 87(12), 1263-1270
SO
     CODEN: JJCREP; ISSN: 0910-5050
PB
     Japanese Cancer Association
     Journal
DT
LA
     English
L12
     ANSWER 112 OF 129 CAPLUS COPYRIGHT 2003 ACS
     1996:742682 CAPLUS
AN
DN
     126:14458
     Expression of multidrug-resistance-associated protein (
ΤI
     MRP) and chemosensitivity in human gastric cancer
AU
     Endo, Kazuya; Maehara, Yoshihiko; Kusumoto, Tetsuya; Ichiyoshi, Yuji;
     Kuwano, Michihiko; Sugimachi, Keizo
CS
     Cancer Center Kyushu, University Hospital, Fukuoka, Japan
     International Journal of Cancer (1996), 68(3), 372-377
SO
     CODEN: IJCNAW; ISSN: 0020-7136
PΒ
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DT
     Journal
     English ·
LA
L12
     ANSWER 113 OF 129 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1996:646439 CAPLUS
DN
     125:266006
ΤI
     Use of protein kinase inhibitors in preventing multidrug
     resistance in cancer cells
IN
     Chaudhary, Preet; Shtil, Alexander A.; Roninson, Igor B.
PΑ
     Board of Trustees of the University of Illinois, USA
SO
     PCT Int. Appl., 75 pp.
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DT
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LΑ
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     WO 1996-US422
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L12 ANSWER 114 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1996:637176 CAPLUS
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     125:266007
     Human vault ribonucleoprotein major protein cDNA sequence and
TI
     vault-related multidrug resistant cancer cell identification using nucleic
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acid or antibody

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ΙN
     Scheper, Riekeld Johannes; Scheffer, George Lodewijk
PA
     Akzo Nobel N.V., Neth.
SO
     PCT Int. Appl., 105 pp.
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PRAI EP 1995-200543
                            19950306
     WO 1996-EP1013
                            19960306
L12 ANSWER 115 OF 129 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1996:546481 CAPLUS
DN
     125:218034
TΙ
     Characterization of the ATP-dependent LTC4 transporter in
     cisplatin-resistant human KB cells
AU
     Chuman, Yutaka; Chen, Zhe-Sheng; Sumizawa, Tomoyuki; Furukawa, Tatsuhiko;
     Haraguchi, Misako; Takebayashi, Yuji; Niwa, Kiyoshi; Yamada, Kazutaka;
     Aikou, Takashi; Akiyama, Shin-ichi
CS
     Inst. Cancer Res., Kagoshima Univ., Kagoshima, Japan
     Biochemical and Biophysical Research Communications (1996), 226(1),
SO
     158-165
     CODEN: BBRCA9; ISSN: 0006-291X
PB · Academic
DT
     Journal
LΑ
     English
L12 ANSWER 116 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1996:419834 CAPLUS
DN
     125:111505
     MRP is frequently expressed in human lung-cancer cell lines, in
TI
     non-small-cell lung cancer and in normal lung
     Giaccone, Giuseppe; Van Ark-Otte, Jannette; Rubio, Gonzalo J.; Gazdar, Adi
ΑU
     F.; Broxterman, Henk J.; Dingemans, Anne-Marie C.; Flens, Marcel J.;
     Scheper, Rik J.; Pinedo, Herbert M.
CS
     University Hospital, Vrije Universiteit, Amsterdam, 1007 MB, Neth.
SO
     International Journal of Cancer (1996), 66(6), 760-767
     CODEN: IJCNAW; ISSN: 0020-7136
PΒ
     Wiley-Liss
DT
     Journal
LA
     English
     ANSWER 117 OF 129 CAPLUS COPYRIGHT 2003 ACS
L12
ΑN
     1996:381566 CAPLUS
DN
     125:51101
     Coordinated induction of MRP/GS-X pump and .gamma.-
TI
     glutamylcysteine synthetase by heavy metals in human leukemia cells
ΑU
     Ishikawa, Toshihisa; Bao, Jia-Ju; Yamane, Yoshiaki; Akimaru, Kunihiro;
     Frindrich, Karl; Wright, Christine D.; Kuo, M. Tien
     Section Molecular Therapeutics, University Texas M. D. Anderson Cancer
CS
     Center, Houston, TX, 77030, USA
     Journal of Biological Chemistry (1996), 271(25), 14981-14988
SO
     CODEN: JBCHA3; ISSN: 0021-9258
     American Society for Biochemistry and Molecular Biology
PB
DT
     Journal
LΑ
     English
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1.12
     ANSWER 118 OF 129 CAPLUS COPYRIGHT 2003 ACS
     1996:305769 CAPLUS
AN
DN
     125:586
TΙ
     Mechanisms of resistance of human small cell lung cancer lines
     selected in VP-16 and cisplatin
     Jain, Nidhi; Lam, Yuk-Miu; Pym, John; Campling, Barbara G.
ΑU
CS
     Cancer Research Laboratories, Queen's University, Kingston, Can.
SO
     Cancer (New York) (1996), 77(9), 1797-1808
     CODEN: CANCAR; ISSN: 0008-543X
PB
     Wiley-Liss
DT
     Journal
LA
     English
     ANSWER 119 OF 129 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1996:261889 CAPLUS
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     124:339679
TI
     Multidrug resistance-associated protein expression in clinical
     gastric carcinoma
ΑU
     Endo, Kazuya; Maehara, Yoshihiko; Ichiyoshi, Yuji; Kusumoto, Tetsuya;
     Sakaguchi, Yoshihisa; Ohno, Shinji; Suqimachi, Keizo
CS
     Cancer Center, Kyushu University Hospital, Fukuoka, 812, Japan
SO
     Cancer (New York) (1996), 77(8, Suppl.), 1681-7
     CODEN: CANCAR; ISSN: 0008-543X
PB
     Wiley-Liss
DT
     Journal
LΑ
     English
L12 ANSWER 120 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1996:228317 CAPLUS
DN
     124:285652
ΤI
     Alterations in expression of the multidrug resistance-associated
     protein (MRP) gene in high-grade transitional cell carcinoma of
     the bladder
ΑU
     Clifford, S. C.; Neal, D. E.; Lunec, J.
CS
     Medical School, University of Newcastle-upon-Tyne, Newcastle-upon-Tyne,
     NE2 4HH, UK
SO
     British Journal of Cancer (1996), 73(5), 659-66
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DΤ
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LΑ
     English
     ANSWER 121 OF 129 CAPLUS COPYRIGHT 2003 ACS
L12
AN
     1996:96400 CAPLUS
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     124:193617
     Non-P-glycoprotein-mediated atypical multidrug resistance in a
ΤI
     human bladder cancer cell line
ΑU
     Naito, Seiji; Hasegawa, Shuji; Yokomizo, Akira; Koga, Hirofumi; Kotoh,
     Shuji; Kuwano, Michihiko; Kumazawa, Joichi
CS
     Fac. Medicine, Kyushu Univ., Fukuoka, 812, Japan
     Japanese Journal of Cancer Research (1995), 86(11), 1112-18
SO
     CODEN: JJCREP; ISSN: 0910-5050
PB
     Japanese Cancer Association
\mathsf{DT}
     Journal
LΑ
     English
L12
    ANSWER 122 OF 129 CAPLUS
                                COPYRIGHT 2003 ACS
ΑN
     1996:51800 CAPLUS
DN
     124:135026
ΤI
     Establishment of a docetaxel-resistant human non-small cell lung cancer
ΑU
     Funayama, Y.; Ohta, S.; Kubota, N.; Nishio, K.; Arioka, H.; Ogasawara, H.;
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- Ohira, T.; Kanazawa, F.; Hasegawa, S.; Saijo, N.
- CS Pharmacol. Div., Natl. Cancer Center Research Inst., Tokyo, 104, Japan
- SO Cellular Pharmacology (1995), 2(6), 303-9 CODEN: CEPHEG; ISSN: 1351-3214
- PB Stockton
- DT Journal
- LA English
- L12 ANSWER 123 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1995:942909 CAPLUS
- DN 124:402
- TI Expression of multidrug resistance-associated protein in NIH/3T3 cells confers multidrug resistance associated with increased drug efflux and altered intracellular drug distribution
- AU Breuninger, Lisa M.; Paul, Saptarshi; Gaughan, Kathleen; Miki, Toru; Chan, Andrew; Aaronson, Stuart A.; Kruh, Gary D.
- CS Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA
- SO Cancer Research (1995), 55(22), 5342-7 CODEN: CNREA8; ISSN: 0008-5472
- PB American Association for Cancer Research
- DT Journal
- LA English
- L12 ANSWER 124 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1995:724074 CAPLUS
- DN 123:160208
- TI MRP gene overexpression in a human doxorubicin-resistant SCLC cell line: alterations in cellular pharmacokinetics and in pattern of cross-resistance
- AU Binaschi, Monica; Supino, Rosanna; Gambetta, Romolo A.; Giaccone, Giuseppe; Prosperi, Ennio; Capranico, Giovanni; Cataldo, Ignazio; Zunino, Franco
- CS Division of Experimental Oncology B, Istituto Nazionale Tumori, Milan, 20133, Italy
- SO International Journal of Cancer (1995), 62(1), 84-9 CODEN: IJCNAW; ISSN: 0020-7136
- DT Journal
- LA English
- L12 ANSWER 125 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1995:670498 CAPLUS
- DN 123:102201
- TI Drug resistance mechanisms and MRP expression in response to epirubicin treatment in a human leukemia cell line
- AU Davey, Ross A.; Longhurst, Terry J.; Davey, Mary W.; Belov, Larissa; Harvie, Rozelle M.; Hancox, Djemilla; Wheeler, Helen
- CS Department Clinical Oncology, Royal North Shore Hospital, St. Leonards, 2065, Australia
- SO Leukemia Research (1995), 19(4), 275-82 CODEN: LEREDD; ISSN: 0145-2126
- PB Elsevier
- DT Journal
- LA English
- L12 ANSWER 126 OF 129 CAPLUS COPYRIGHT 2003 ACS
- AN 1995:535813 CAPLUS
- DN 122:305991
- TI The role of methoxymorpholino anthracycline and cyanomorpholino anthracycline in a sensitive small-cell lung-cancer cell line and its multidrug-resistant but P-glycoprotein-negative and cisplatin-resistant counterparts
- AU Graaf, Winette T. A. van der; Mulder, Nanno H.; Meijer, Coby; Vries,

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Elisabeth G. E. de
CS
     Department Internal Medicine, University Hospital, Groningen, 9713 EZ,
     Cancer Chemotherapy and Pharmacology (1995), 35(4), 345-8
SO
     CODEN: CCPHDZ; ISSN: 0344-5704
DT
     Journal
LΑ
     English
L12
     ANSWER 127 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1995:209914 CAPLUS
     122:527
DN
ΤI
     Pharmacological characterization of multidrug resistant MRP
     -transfected human tumor cells
· AU
     Cole, Susan P. C.; Sparks, Kathryn E.; Fraser, Karen; Loe, Douglas W.;
     Grant, Caroline E.; Wilson, Gerald M.; Deeley, Roger G.
     Cancer Res. Lab., Queen's Univ., Kingston, ON, K7L 3N6, Can.
CS
SO
     Cancer Research (1994), 54(22), 5902-10
     CODEN: CNREA8; ISSN: 0008-5472
     American Association for Cancer Research
PB
DT
     Journal
LΑ
     English
L12 ANSWER 128 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1994:595251 CAPLUS
DN
     121:195251
TI
     Prevalence of multidrug resistance related to activation of the
     mdrl gene in human sarcoma mutants derived by single-step doxorubicin
     selection
ΑU
     Chen, Gang; Jaffrezou, Jean Pierre; Fleming, William H.; Duran, George E.;
     Sikic, Branimir I.
CS
     Sch. Med., Stanford Univ., Stanford, CA, 94305-5306, USA
SO
     Cancer Research (1994), 54(18), 4980-7
     CODEN: CNREA8; ISSN: 0008-5472
DT
     Journal
LΑ
     English
L12
    ANSWER 129 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1994:94935 CAPLUS
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     120:94935
TΤ
     Cross-resistance to diverse drugs is associated with primary
     cisplatin resistance in ovarian cancer cell lines
     Hamaguchi, Kinya; Godwin, Andrew K.; Yakushiji, Michiaki; O'Dwyer, Peter
ΑU
     J.; Ozols, Robert F.; Hamilton, Thomas C.
     Dep. Med. Oncol., Fox Chase Cancer Cent., Philadelphia, PA, 19111, USA
CS-
     Cancer Research (1993), 53(21), 5225-32
SO
     CODEN: CNREA8; ISSN: 0008-5472
DT
     Journal
LA
     English
=> d 112 113 all
    ANSWER 113 OF 129 CAPLUS COPYRIGHT 2003 ACS
     1996:646439 CAPLUS
AN
DN
     125:266006
     Use of protein kinase inhibitors in preventing multidrug
TΤ
     resistance in cancer cells
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Chaudhary, Preet; Shtil, Alexander A.; Roninson, Igor B.

Board of Trustees of the University of Illinois, USA

IN

PΑ

SO

DT

PCT Int. Appl., 75 pp.

CODEN: PIXXD2

Patent

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English
LA
IC
     ICM A61K045-06
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AB
     Methods are disclosed for preventing the emergence of multidrug
     resistance in tumor cells during cancer chemotherapy. In
     particular, protein kinase inhibitors are used to prevent the induction of
     expression of the multidrug resistance gene (MDR1) encoding
     P-glycoprotein by chemotherapeutic drugs. MDR1 expression, which results
     in tumor cell resistance to subsequent treatment with certain
     chemotherapeutic drugs, is shown herein to be induced in response to
     treatment with various cytotoxic agents, including such agents that are
     and are not substrates for P-glycoprotein-mediated efflux from cancer
     cells. Inhibitors of protein kinases, in particular protein kinase C, are
     shown to suppress this cellular response. In addn., such protein kinase
     inhibitors are also shown to inhibit expression of a gene encoding a
     multidrug resistance-assocd. protein (the MRP gene).
     Methods are disclosed for using such protein kinase inhibitors to both
     suppress induction of MDR1 gene expression by cytotoxic drugs and to
     inhibit expression of MRP. Also provided are methods for
     identifying protein kinase inhibitors that have either or both of these
     effects on MDR1 and MRP expression. Thus, the invention
     provides useful methods and reagents for preventing the emergence of
     multidrug resistance in tumor cells treated with cytotoxic and
     chemotherapeutic drugs in cancer patients undergoing chemotherapy, when
     such protein kinase inhibitors are administered prior to or simultaneous
     with cytotoxic drug treatment in such individuals.
     protein kinase inhibitor multidrug resistance inhibition; MDR
ST
     inhibition protein kinase inhibitor; cancer therapy protein kinase
     inhibitor MDR
ΙT
     Gene, animal
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (MRP; protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Lymphocyte
        (differentiation; protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Cell differentiation
        (lymphoid cell; protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Proteins, specific or class, biological studies
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (multidrug resistance-assocd.; protein kinase inhibitors for
        prevention of multidrug resistance in cancer cells)
IT
     Biological transport
     Cytotoxic agents
    HeLa cell
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Lymphoma
     Neoplasm inhibitors
         (protein kinase inhibitors for prevention of multidrug
         resistance in cancer cells)
IT
     Glycophosphoproteins
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
      (Biological study); PROC (Process)
         (P-, gene mdrl, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
         (T-cell, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
         (acute monocytic, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Leukemia
        (acute myelogenous, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
     Uterus, neoplasm
        (cervix, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
ΙT
     Therapeutics
        (chemo-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
TT
     Leukemia
        (chronic myelocytic, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
ΙT
     Skin, neoplasm
        (epidermoid carcinoma, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
        (fibro-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Neoplasm inhibitors
        (hematol., protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Neoplasm inhibitors
        (leukemia, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Gene, animal
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (mdrl, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Drug resistance
        (multi-, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Bladder
     Mammary gland
        (neoplasm, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Thymus gland
        (neoplasm, thymoma, protein kinase inhibitors for prevention of
        multidrug resistance in cancer cells)
IT
     Leukemia
        (promyelocytic, protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
IT
     Neoplasm inhibitors
        (solid, protein kinase inhibitors for prevention of multidrug
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IT
        (white, and calphostin C; protein kinase inhibitors for prevention of
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multidrug resistance in cancer cells)
IT
     59-05-2, Methotrexate 127-07-1, Hydroxyurea 147-94-4, Cytosine
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                   305-03-3, Chlorambucil 446-72-0, Genistein 865-21-4,
                   1405-10-3, Neomycin sulfate 1405-10-3D, Neomycin sulfate,
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     25316-40-9, Adriamycin
                                                           34316-15-9D,
     Chelerythrine, derivs.
                              62996-74-1, Staurosporine
                                                          62996-74-1D,
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                              63177-57-1, Methyl 2,5-dihydroxycinnamate
                                 84477-87-2, H7 84477-87-2D, H7, derivs.
     70563-58-5, Herbimycin A
                 91742-10-8, HA1004 100827-28-9, Erbstatin 100827-28-9D, derivs. 118409-58-8, Tyrphostin A25 118409-58-8D,
     88494-43-3
     Erbstatin, derivs.
     Tyrphostin A25, derivs.
                               121263-19-2, Calphostin C
                                                            121263-19-2D,
     Calphostin C, derivs.
                             149092-34-2, Tyrphostin B46
                                                            149092-34-2D,
     Tyrphostin B46, derivs.
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (protein kinase inhibitors for prevention of multidrug
        resistance in cancer cells)
TΤ
     9026-43-1, Protein kinase
                                 141436-78-4, Protein kinase C
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
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        resistance in cancer cells)
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L12 ANSWER 120 OF 129 CAPLUS COPYRIGHT 2003 ACS
AN
     1996:228317 CAPLUS
DN
     124:285652
     Alterations in expression of the multidrug resistance-associated
     protein (MRP) gene in high-grade transitional cell carcinoma of
     the bladder
ΑU
     Clifford, S. C.; Neal, D. E.; Lunec, J.
     Medical School, University of Newcastle-upon-Tyne, Newcastle-upon-Tyne,
CS
     NE2 4HH, UK
SO
     British Journal of Cancer (1996), 73(5), 659-66
     CODEN: BJCAAI; ISSN: 0007-0920
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     Stockton
DT
     Journal
LA
     English
     14-1 (Mammalian Pathological Biochemistry)
     Section cross-reference(s): 1
     Expression of the MRP gene has been demonstrated in vitro to be
AΒ
     a causal factor in non-P-glycoprotein-mediated multidrug
     resistance, and is implicated in resistance to a no. of
     the chemotherapeutic agents currently used in the treatment of high-grade
     transitional cell carcinoma (TCC) of the bladder (doxorubicin, epirubicin
     and vinblastine). Using a sensitive RT-PCR-based technique, we have
     quantified MRP mRNA levels in a series of untreated TCC (n=24),
     normal bladder (n=5) and control tissue and cell line samples.
     MRP mRNA was widely expressed and detectable in all samples
     analyzed, with considerable (up to 190-fold) variation obsd. between
     individual tumor samples. MRP mRNA levels found in TCC samples
     were lower than those detd. for normal peripheral mononucleocyte
     (2.3-fold) and testis (4.1-fold) samples, previously reported to be
     high-expressing tissues, and varied over a similar range to that obsd. in
     normal bladder samples. Results indicate that MRP mRNA levels
    in a greater proportion of high-grade (G3) bladder tumors (55%, 6/11) are
     significantly reduced (P=0.018) compared with low- and moderate-grade
     (G1/2) bladder tumors (8%, 1/13), and suggest that MRP mRNA
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levels frequently become reduced as a consequence of tumor progression to
     advanced, poorly differentiated disease. No correlation was apparent
     between MRP and MDR1 mRNA levels, thus providing no evidence to
     suggest common regulation of the two genes. In a limited no. of patients,
     no evidence was found to support a role for MRP mRNA levels as a
     determinant of response to chemotherapy in patients being uniformly
     treated with either cisplatin-methotrexate-vinblastine (n=6) or
     epirubicin-cisplatin-methotrexate (n=4) regimens. Similarly, no overall
     pattern of altered MRP mRNA expression was obsd. following
     chemotherapy in four patients from whom post chemotherapy biopsies were
     taken. This study provides a useful pilot investigation regarding the
     level, variation and pattern of MRP mRNA expression in TCC of
    the bladder, and suggests that further studies to establish the clin.
     significance of these variations are required.
     multidrug resistance protein gene bladder carcinoma
     Gene, animal
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (MRP; multidrug resistance-assocd. protein gene
        expression in human high-grade transitional cell carcinoma of the
     Proteins, specific or class
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (multidrug resistance-assocd. protein (MRP);
        multidrug resistance-assocd. protein gene expression in human
        high-grade transitional cell carcinoma of the bladder)
     Neoplasm inhibitors
        (multidrug resistance-assocd. protein gene expression in
        human high-grade transitional cell carcinoma of the bladder)
     Ribonucleic acids, messenger
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (multidrug resistance-assocd. protein gene expression in
        human high-grade transitional cell carcinoma of the bladder)
     Bladder
        (neoplasm, transitional cell carcinoma, multidrug resistance
        -assocd. protein gene expression in human high-grade transitional cell
        carcinoma of the bladder)
     59-05-2, Methotrexate
                             865-21-4, Vinblastine 15663-27-1,
     Cisplatin
                56420-45-2, Epirubicin
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (multidrug resistance-assocd. protein gene expression in
        human high-grade transitional cell carcinoma of the bladder)
=> d 112 121 all
    ANSWER 121 OF 129 CAPLUS. COPYRIGHT 2003 ACS
     1996:96400 CAPLUS
    124:193617
    Non-P-glycoprotein-mediated atypical multidrug resistance in a
    human bladder cancer cell line
    Naito, Seiji; Hasegawa, Shuji; Yokomizo, Akira; Koga, Hirofumi; Kotoh,
    Shuji; Kuwano, Michihiko; Kumazawa, Joichi
    Fac. Medicine, Kyushu Univ., Fukuoka, 812, Japan
    Japanese Journal of Cancer Research (1995), 86(11), 1112-18
    CODEN: JJCREP; ISSN: 0910-5050
    Japanese Cancer Association
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L6 53 L3

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L7 2 L5 AND L6

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- L7 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS
- AN 2003:3461 CAPLUS
- TI Breast cancer resistance protein (BCRP/ABCG2) induces cellular resistance to HIV-1 nucleoside reverse transcriptase inhibitors
- AU Wang, Xin; Furukawa, Tatsuhiko; Nitanda, Takao; Okamoto, Mika; Sugimoto, Yoshikazu; Akiyama, Shin-Ichi; Baba, Masanori
- CS Division of Human Retroviruses, Center for Chronic Viral Diseases, Faculty of Medicine, Kagoshima University, Kagoshima, Japan
- SO Molecular Pharmacology (2003), 63(1), 65-72 CODEN: MOPMA3; ISSN: 0026-895X
- PB American Society for Pharmacology and Experimental Therapeutics
- DT Journal
- LA English
- CC 1-5 (Pharmacology)
- AΒ Breast cancer resistance protein (BCRP/ABCG2) is a novel member of ATP-binding cassette transporters, which induce multidrug resistance in cancer cells. We found that a high level of BCRP expression in CD4+ T cells conferred cellular resistance to human immunodeficiency virus type-1 (HIV-1) nucleoside reverse transcriptase inhibitors. The cell line MT-4/DOX500 was established through the long-term culture of MT-4 cells in the presence of doxorubicin (DOX) and had reduced sensitivity to not only DOX but also zidovudine (AZT). MT-4/DOX500 cells showed reduced intracellular accumulation and retention of DOX and increased ATP-dependent rhodamine 123 efflux. The cells were also resistant to several anticancer agents such as mitoxantrone, 7-ethyl-10-[4-(1piperidino)-1-piperidino]carbonyloxycamptothecin, and 7-ethyl-10hydroxycamptothecin. AZT was 7.5-fold less inhibitory to HIV-1 replication in MT-4/DOX500 cells than in MT-4 cells. Furthermore, the anti-HIV-1 activity of lamivudine was severely impaired in MT-4/DOX500 cells. In contrast, the antiviral activity of non-nucleoside reverse transcriptase inhibitors and protease inhibitors was not affected in the cells. MT-4/DOX500 cells expressed glycosylated BCRP but not P-glycoprotein (ABCB1), multidrug resistance protein 1, 2, or 4 (ABCC1, -2, or -4), or lung resistance-related protein. In addn., the BCRP-specific inhibitor fumitremorgin C completely abolished the resistance of MT-4/DOX500 cells to AZT as well as to DOX. An anal. for intracellular metab. of AZT suggests that the resistance is attributed to the increase of ATP-dependent efflux of its metabolites, presumably AZT 5'-monophosphate, in MT-4/DOX500 cells.
- ST lamivudine zidovudine antiviral BCRP ABCG2 protein anticancer doxorubicin resistance
- IT Multidrug resistance proteins
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (BCRP (breast cancer resistance protein); breast cancer resistance
 protein induces cellular resistance to HIV-1 NRTIs in DOX-resistant
 CD4+ T-cell lines)
- IT Multidrug resistance proteins
 - RL: BSU (Biological study, unclassified); BIOL (Biological study) (MRP1; breast cancer resistance protein induces cellular resistance to HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
- IT Multidrug resistance proteins
 - RL: BSU (Biological study, unclassified); BIOL (Biological study) (MRP2; breast cancer resistance protein induces cellular resistance to

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HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     Multidrug resistance proteins
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
         (MRP4; breast cancer resistance protein induces cellular resistance to
        HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     Drug resistance
         (antitumor; breast cancer resistance protein induces cellular
        resistance to HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     Glycosylation
         (biol.; breast cancer resistance protein induces cellular resistance to
        HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     Antitumor agents
     Antiviral agents
     Cytotoxic agents
     Human
     Multidrug resistance
         (breast cancer resistance protein induces cellular resistance to HIV-1
        NRTIs in DOX-resistant CD4+ T-cell lines)
IΤ
     Antitumor agents
         (resistance to; breast cancer resistance protein induces cellular
        resistance to HIV-1 NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     118974-02-0, Fumitremorgin C
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
         (breast cancer resistance protein induces cellular resistance to HIV-1
        NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     23214-92-8, Doxorubicin 29706-85-2, AZT 5'-monophosphate
     Zidovudine 92586-35-1, 3'-Azido-3'-deoxythymidine triphosphate 106060-89-3, 3'-Azido-3'-deoxythymidine diphosphate
     RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT
     (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
         (breast cancer resistance protein induces cellular resistance to HIV-1
        NRTIs in DOX-resistant CD4+ T-cell lines)
IT
     50-76-0, Actinomycin D
                              57-22-7, Vincristine
                                                       3056-17-5, Stavudine
     15663-27-1, Cisplatin
                              33069-62-4, Paclitaxel
                                                      33419-42-0,
     Etoposide
                 62669-70-9, rhodamine 123
                                               65271-80-9, Mitoxantrone
     69655-05-6, Didanosine
                               100286-90-6, CPT-11
                                                     129618-40-2, Nevirapine
                                149950-60-7, Emivirine
     134678-17-4, Lamivudine
                                                          150378-17-9, Indinavir
     159989-64-7, Nelfinavir
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (breast cancer resistance protein induces cellular resistance to HIV-1
        NRTIs in DOX-resistant CD4+ T-cell lines)
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- L7 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2003 ACS
- AN 2000:72225 CAPLUS
- DN 132:216690
- TI Fumitremorgin C reverses multidrug resistance in cells transfected with the breast cancer resistance protein
- AU Rabindran, Sridhar K.; Ross, Douglas D.; Doyle, L. Austin; Yang, Weidong; Greenberger, Lee M.
- CS Oncology and Immunoinflammatory Research, Wyeth-Ayerst Research, Pearl River, NY, 10965, USA
- SO Cancer Research (2000), 60(1), 47-50 CODEN: CNREA8; ISSN: 0008-5472
- PB AACR Subscription Office
- DT Journal
- LA English
- CC 1-6 (Pharmacology)
- Fumitremorgin C (FTC) is a potent and specific chemosensitizing agent in cell lines selected for resistance to mitoxantrone that do not overexpress P-glycoprotein or multidrug resistance protein. The gene encoding a novel transporter, the breast cancer resistance protein (BCRP), was recently overexpressed in a mitoxantrone-selected human colon cell line, S1-M1-3.2, which was used to identify FTC. Because the drug-selected cell line may contain multiple alterations contributing to the multidrug resistance phenotype, the authors examd. the effect of FTC on MCF-7 cells transfected with the BCRP gene. The authors report that FTC almost completely reverses resistance mediated by BCRP in vitro and is a pharmacol. probe for the expression and mol. action of this transporter.
- ST fumitremorgin C multidrug resistance reversal; breast cancer resistance protein fumitremorgin C
- IT Drug resistance
 - (antitumor; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)
- IT Multidrug resistance proteins
 - RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)
 - (breast cancer resistance protein (BCRP); fumitremorgin C reverses

multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT Biological transport

(drug; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT Multidrug resistance

(fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT Antitumor agents

(mammary gland; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT Mammary gland

Mammary gland

(neoplasm, inhibitors; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT Antitumor agents

(resistance to; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT 20830-81-3, Daunorubicin 23214-92-8, Doxorubicin

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT 57-22-7, Vincristine 15663-27-1, cis-Platinum 33069-62-4,
 Paclitaxel 65271-80-9, Mitoxantrone 118974-02-0, Fumitremorgin
 C 123948-87-8, Topotecan

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

IT 225918-89-8, BBR 3390

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(intracellular accumulation; fumitremorgin C reverses multidrug resistance in cells transfected with breast cancer resistance protein in relation to effect on drug transport)

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L9
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AN
DN
     138:332878
ΤI
     Application of a human multidrug transporter (abcg2) variant as selectable
     marker in gene transfer to progenitor cells and in gene therapy
IN
     Nemet, Katalin; Varadi, Gyorgy; Cervenak, Judit; Ujhelly, Olga; Sarkadi,
     Balazs; Varadi, Andras; Oezvegy, Csilla
PA
     Solvo Biotechnology Inc., Hung.
so
     PCT Int. Appl., 44 pp.
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LA
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FAN.CNT 2
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L9
     ANSWER 2 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     2003:3461 CAPLUS
TI
     Breast cancer resistance protein (BCRP/ABCG2) induces cellular
     resistance to HIV-1 nucleoside reverse transcriptase inhibitors
ΑU
     Wang, Xin; Furukawa, Tatsuhiko; Nitanda, Takao; Okamoto, Mika; Sugimoto,
     Yoshikazu; Akiyama, Shin-Ichi; Baba, Masanori
     Division of Human Retroviruses, Center for Chronic Viral Diseases, Faculty
CS
     of Medicine, Kagoshima University, Kagoshima, Japan
SO
     Molecular Pharmacology (2003), 63(1), 65-72
     CODEN: MOPMA3; ISSN: 0026-895X
     American Society for Pharmacology and Experimental Therapeutics
PB
DT
     Journal
     English
RE.CNT 40
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              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 3 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:696659 CAPLUS
DN
     137:222100
     Improving bioavailability of orally administered drugs, screening for
ΤI
     enhancers of such bioavailability and oral drug delivery compositions
IN
     Schellens, Johannes Henricus Matthias; Schinkel, Alfred Hermanus
     Netherlands Cancer Institute, Neth.
PΑ
SO
     U.S. Pat. Appl. Publ., 11 pp., Cont.-in-part of Appl. No. PCT/NL00/00331.
     CODEN: USXXCO
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     English
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PRAI NL 1999-1012066
                          Α
                                19990517
      NL 1999-1012481
                          Α
                                19990630
      WO 2000-NL331
                                20000517
                           A2
L9
      ANSWER 4 OF 15 CAPLUS COPYRIGHT 2003 ACS
ΑN
      2002:696257 CAPLUS
DN
      137:226574
ΤI
      Screening system based on expression of ABCG2 half transporter protein
IN
      Oezvegy, Csilla; Szakacs, Gergely; Varadi, Andras; Nagy, Zoltan
PA
      Solvo Biotechnology Inc., Hung.
      PCT Int. Appl., 26 pp.
SO
      CODEN: PIXXD2
DT
      Patent
LΑ
      English
FAN.CNT 2
      PATENT NO.
                         KIND
                                DATE
                                                  APPLICATION NO.
                                                                     DATE
                                                  -----
ΡI
      WO 2002071073
                          A2
                                20020912
                                                  WO 2002-HU15
                                                                     20020304
      WO 2002071073
                          A3
                                20030403
              AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
               CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
               GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
               LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
               PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
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               TJ, TM
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      WO 2003035685
                          A1
                               20030501
                                                WO 2002-HU108
                                                                  20021024
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PRAI HU 2001-947
                          Α
                                20010302
     HU 2001-4446
                          Α
                                20011024
     WO 2002-HU15
                          Α
                                20020304
     HU 2002-3435
                          Α
                                20021011
L9
     ANSWER 5 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:524344 CAPLUS
DN
     138:201203
TΙ
     Flow cytometric analysis of breast cancer resistance protein
     expression and function
```

- AU Minderman, Hans; Suvannasankha, Attaya; O'Loughlin, Kieran L.; Scheffer, George L.; Scheper, Rik J.; Robey, Robert W.; Baer, Maria R.
- CS Leukemia Section, Department of Medicine, Roswell Park Cancer Institute, Buffalo, NY, USA
- SO Cytometry (2002), 48(2), 59-65 CODEN: CYTODQ; ISSN: 0196-4763
- PB Wiley-Liss, Inc.
- DT Journal
- LA English
- RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L9 ANSWER 6 OF 15 CAPLUS COPYRIGHT 2003 ACS
- AN 2002:290794 CAPLUS
- DN 136:304053
- TI Reversal of multidrug **resistance** in human colon carcinoma cells using fumitremorgins and diketopiperazines
- IN Rabindran, Sridhar Krishna; He, Haiyin; Greenberger, Lee Martin
- PA American Cyanamid Company, USA
- SO U.S., 19 pp. CODEN: USXXAM
- DT Patent
- LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. DATE | | | |
|------|-----------------|------|----------|--------------------------|--|--|--|
| | | | | | | | |
| ΡI | บร 6372775 | В1 | 20020416 | US 1999-321182 19990527 | | | |
| | US 2002156015 | A1 | 20021024 | US 2002-86169 20020228 | | | |
| | US 2002169111 | A1 | 20021114 | . US 2002-86132 20020228 | | | |
| | US 6537964 | B1 | 20030325 | US 2002-86170 20020228 | | | |
| | US 2003083230 | A1 | 20030501 | US 2002-86133 20020228 | | | |
| PRAI | US 1998-109801P | P | 19980527 | | | | |
| | US 1999-321182 | A3 | 19990527 | | | | |
| ~ ~ | 100000 100 0040 | | | | | | |

OS MARPAT 136:304053

RE.CNT 72 THERE ARE 72 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L9 ANSWER 7 OF 15 CAPLUS COPYRIGHT 2003 ACS
- AN 2001:487684 CAPLUS
- DN 135:328480
- TI Functional Characterization of the Human Multidrug Transporter, ABCG2, Expressed in Insect Cells
- AU Ozvegy, Csilla; Litman, Thomas; Szakacs, Gergely; Nagy, Zoltan; Bates, Susan; Varadi, Andras; Sarkadi, Balazs
- CS Institute of Enzymology, Biological Research Center, Hungarian Academy of Sciences, Budapest, H-1113, Hung.
- SO Biochemical and Biophysical Research Communications (2001), 285(1), 111-117
 - CODEN: BBRCA9; ISSN: 0006-291X
- PB Academic Press
- DT Journal
- LA English
- RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L9 ANSWER 8 OF 15 CAPLUS COPYRIGHT 2003 ACS
- AN 2001:429174 CAPLUS
- DN 135:285265
- TI A functional assay for detection of the mitoxantrone resistance protein, MXR (ABCG2)
- AU Robey, R. W.; Honjo, Y.; van de Laar, A.; Miyake, K.; Regis, J. T.; Litman, T.; Bates, S. E.

```
CS
      Center for Cancer Research, Medicine Branch, Developmental Therapeutics
     Department, National Cancer Institute, National Institutes of Health,
      Bethesda, MD, 20892, USA
      Biochimica et Biophysica Acta (2001), 1512(2), 171-182
      CODEN: BBACAQ; ISSN: 0006-3002
PB
     Elsevier Science B.V.
DT
      Journal
LΑ
     English
RE.CNT 40
               THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 9 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
      2000:872654 CAPLUS
DN
     134:216800
TI
     Inhibition of BCRP-mediated drug efflux by fumitremorgin-type indolyl
     diketopiperazines
     van Loevezijn, A.; Allen, J. D.; Schinkel, A. H.; Koomen, G.-J.
ΑU
     Institute of Molecular Chemistry, Laboratory of Organic Chemistry,
CS
     University of Amsterdam, Amsterdam, NL-1018 WS, Neth.
     Bioorganic & Medicinal Chemistry Letters (2000), Volume Date 2001, 11(1),
SO
     CODEN: BMCLE8; ISSN: 0960-894X
PΒ
     Elsevier Science Ltd.
DT
     Journal
     English
LΑ
RE.CNT 23
               THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 10 OF 15 CAPLUS COPYRIGHT 2003 ACS
ΑN
     2000:824069 CAPLUS
DN
     134:9341
     A method of improving bioavailability of orally administered drugs,
TI
     screening for enhancers of such bioavailability and novel pharmaceutical
     compositions for oral delivery of drugs
IN
     Schellens, Johannes Henricus Matthias; Schinkel, Alfred Hermanus
PA
     Het Nederlands Kankerinstituut, Neth.
SO
     PCT Int. Appl., 25 pp.
     CODEN: PIXXD2
DT
     Patent
     English
FAN.CNT 2
     PATENT NO.
                        KIND
                              DATE
                                               APPLICATION NO.
                                                                  DATE
                        ____
                              _____
                                               ______
PΙ
     WO 2000069390
                         A2
                               20001123
                                               WO 2000-NL331
                                                                  20000517
     WO 2000069390
                        A3
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     AU 2000049552
                         A5
                             20010205
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                                                                  20000517
     EP 1189637
                         A2
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             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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                         Α1
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                                                                  20011119
PRAI NL 1999-1012066
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                         Α
     NL 1999-1012481
                         Α
                               19990630
     WO 2000-NL331
                         W
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L9
     ANSWER 11 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     2000:72225 CAPLUS
DN
     132:216690
TI
     Fumitremorgin C reverses multidrug resistance in cells
     transfected with the breast cancer resistance protein
ΑU
     Rabindran, Sridhar K.; Ross, Douglas D.; Doyle, L. Austin; Yang, Weidong;
     Greenberger, Lee M.
     Oncology and Immunoinflammatory Research, Wyeth-Ayerst Research, Pearl
CS
     River, NY, 10965, USA
SO
     Cancer Research (2000), 60(1), 47-50
     CODEN: CNREA8; ISSN: 0008-5472
PB
     AACR Subscription Office
DT
     Journal
LA
     English
RE.CNT 20
              THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 12 OF 15 CAPLUS COPYRIGHT 2003 ACS
AN
     2000:27209 CAPLUS
DN
     132:180759
     Fumitremorgin C analogs that reverse mitoxantrone resistance in
     human colon carcinoma cells
ΑU
     He, Haiyin; Rabindran, Sridhar G.; Greenberger, Lee M.; Carter, Guy T.
CS
     Natural Products Chemistry, Wyeth-Ayerst Research, Pearl River, NY, 10965,
SO
     Medicinal Chemistry Research (1999), 9(6), 424-437
     CODEN: MCREEB; ISSN: 1054-2523
PB
     Birkhaeuser Boston
DT
     Journal
LA
     English
RE.CNT 16
              THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 13 OF 15 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1999:511178 CAPLUS
DN
     131:143080
ΤI
     A multidrug resistance protein associated with antitumor drug
     resistance in breast cancer and a cDNA encoding it
IN
     Ross, Douglas D.; Doyle, L. Austin; Abruzzo, Lynne
PA
     University of Maryland, Baltimore, USA
     PCT Int. Appl., 79 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
FAN.CNT 1
     PATENT NO.
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                            DATE
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                                                             DATE
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PΙ
     WO 9940110
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                                           WO 1999-US2577
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             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
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             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    CA 2319715
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                                                             19990205
    AU 9927610
                       Α1
                            19990823
                                           AU 1999-27610
                                                             19990205
    AU 755567
                       В2
                            20021212
    EP 1054894
                       A1
                            20001129
                                           EP 1999-908097
                                                             19990205
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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"HELP COMMANDS" at an arrow prompt (=>).

=> s e4

L2 16 CISPLATIN/BI

=> s fumitremorgin c

36 FUMITREMORGIN

2061134 C

L3 8 FUMITREMORGIN C

(FUMITREMORGIN(W)C)

=> file caplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
18.08
21.84

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE

0.00 -0.65

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FILE COVERS 1907 - 19 Jun 2003 VOL 138 ISS 25 FILE LAST UPDATED: 18 Jun 2003 (20030618/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

(FILE 'HOME' ENTERED AT 15:54:12 ON 19 JUN 2003)

FILE 'CAPLUS' ENTERED AT 15:54:21 ON 19 JUN 2003

FILE 'REGISTRY' ENTERED AT 15:55:33 ON 19 JUN 2003

L1 0 S CISPLATEN

E CISPLATEN

L2 16 S E4

L3 8 S FUMITREMORGIN C

FILE 'CAPLUS' ENTERED AT 15:57:27 ON 19 JUN 2003

=> s 11

L4 0 L1

=> s 12

L5 14128 L2

111427-99-7, TR 3 111468-06-5 115589-18-9 118974-02-0, Fumitremorgin C 137494-04-3 140715-85-1, Fumiquinazoline A 140715-86-2, Fumiquinazoline D 140715-87-3, Fumiquinazoline E 140852-71-7, Fumiquinazoline B 140924-01-2, Fumiquinazoline C RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (structure and cytotoxic activity of, from Aspergillus fumigatus)

=> file reg COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 3.55 3.76 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE. -0.65-0.65

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 18 JUN 2003 HIGHEST RN 533863-98-8 DICTIONARY FILE UPDATES: 18 JUN 2003 HIGHEST RN 533863-98-8

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> s cisplaten

L1 0 CISPLATEN

=> e cisplaten E11 CISPERMETHRIN/BI E2 1 CISPLAT/BI E3 0 --> CISPLATEN/BI F.4 16 CISPLATIN/BI E5 1 CISPLATINUM/BI E6 1 CISPLATYL/BI E7 5 CISS/BI 2 E8 CISSAGLABERR/BI E9 2 CISSAGLABERRIMINE/BI E10 2 CISSAM/BI E11 5 CISSAMINE/BI 5 E12 CISSAMPAREINE/BI

=> se4

SE4 IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter

```
Welcome to STN International! Enter x:x
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LOGINID:sssptau125rxt

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PASSWORD:
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TERMINAL (ENTER 1, 2, 3, OR ?):2

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Welcome to STN International
NEWS
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2
                 "Ask CAS" for self-help around the clock
NEWS 3
                 New e-mail delivery for search results now available
         Jun 03
NEWS 4
         Aug 08
                 PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 5
         Aug 19 Aquatic Toxicity Information Retrieval (AQUIRE)
                 now available on STN
                 Sequence searching in REGISTRY enhanced
NEWS
         Aug 26
NEWS 7
         Sep 03
                 JAPIO has been reloaded and enhanced
NEWS 8
         Sep 16 Experimental properties added to the REGISTRY file
         Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 9
NEWS 10 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
NEWS 11 Oct 24 BEILSTEIN adds new search fields
NEWS 12 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 13 Nov 18 DKILIT has been renamed APOLLIT
NEWS 14 Nov 25 More calculated properties added to REGISTRY
NEWS 15 Dec 04
                CSA files on STN
NEWS 16 Dec 17
                 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 17
         Dec 17
                 TOXCENTER enhanced with additional content
NEWS 18
                 Adis Clinical Trials Insight now available on STN
         Dec 17
NEWS 19
         Jan 29
                 Simultaneous left and right truncation added to COMPENDEX,
                 ENERGY, INSPEC
NEWS 20 Feb 13
                CANCERLIT is no longer being updated
NEWS 21 Feb 24 METADEX enhancements
NEWS 22 Feb 24 PCTGEN now available on STN
NEWS 23 Feb 24 TEMA now available on STN
NEWS 24 Feb 26 NTIS now allows simultaneous left and right truncation
NEWS 25 Feb 26 PCTFULL now contains images
NEWS 26 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 27 Mar 20 EVENTLINE will be removed from STN
NEWS 28 Mar 24 PATDPAFULL now available on STN
NEWS 29 Mar 24 Additional information for trade-named substances without
                 structures available in REGISTRY
NEWS 30 Apr 11
                Display formats in DGENE enhanced
NEWS 31 Apr 14
                MEDLINE Reload
NEWS 32 Apr 17
                Polymer searching in REGISTRY enhanced
NEWS 33
        Jun 13
                Indexing from 1947 to 1956 added to records in CA/CAPLUS
NEWS 34
        Apr 21
                New current-awareness alert (SDI) frequency in
                WPIDS/WPINDEX/WPIX
NEWS 35
                RDISCLOSURE now available on STN
        Apr 28
NEWS 36 May 05
                Pharmacokinetic information and systematic chemical names
                added to PHAR
NEWS 37
        May 15
                MEDLINE file segment of TOXCENTER reloaded
NEWS 38
        May 15
                Supporter information for ENCOMPPAT and ENCOMPLIT updated
NEWS 39
        May 16 CHEMREACT will be removed from STN
NEWS 40 May 19
                Simultaneous left and right truncation added to WSCA
                RAPRA enhanced with new search field, simultaneous left and
NEWS 41
        May 19
                right truncation
NEWS 42
         Jun 06
                Simultaneous left and right truncation added to CBNB
        Jun 06 PASCAL enhanced with additional data
```

NEWS EXPRESS April 4 CURRENT WINDOWS VERSION IS V6.01a, CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),

AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003

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NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

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=> file caplus COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

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FILE COVERS 1907 - 19 Jun 2003 VOL 138 ISS 25 FILE LAST UPDATED: 18 Jun 2003 (20030618/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d 116:210833 all

ANSWER 1 CAPLUS COPYRIGHT 2003 ACS

AN 1992:210833 CAPLUS

DN 116:210833

TI Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish

AU Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro

CS Osaka Univ. Pharm. Sci., Osaka, Japan

```
SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS

DT Journal

LA Japanese

CC 10-1 (Microbial, Algal, and Fungal Biochemistry)

Section cross-reference(s): 1
```

GI

AB Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior.

ST Aspergillus fumiquinazoline cytotoxicity structure

IT Nomenclature, new natural products

(fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Aspergillus fumigatus

(fumiquinazolines from, structure and cytotoxicity of)

IT Molecular structure, natural product

(of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)

IT Molecular structure, natural product

(of fumiquinazoline E (quinazoline), from Aspergillus fumigatus)

IT Neoplasm inhibitors

ΙT

(leukemia, fumiquinazolines as, from Aspergillus fumigatus)

140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P

IT 67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. 12771-72-1, Verruculogen 51177-07-2 62867-47-4, Fumigaclavine C 74149-38-5

10/086,133

Welcome to STN International! Enter x:x

LOGINID:sssptau125rxt

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
Welcome to STN International
NEWS
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2
                 "Ask CAS" for self-help around the clock
NEWS 3
                 New e-mail delivery for search results now available
         Jun 03
NEWS 4
                 PHARMAMarketLetter(PHARMAML) - new on STN
         Aug 19 Aquatic Toxicity Information Retrieval (AQUIRE)
NEWS 5
                 now available on STN
NEWS
         Aug 26
                 Sequence searching in REGISTRY enhanced
NEWS
         Sep 03
                 JAPIO has been reloaded and enhanced
         Sep 16
NEWS
                 Experimental properties added to the REGISTRY file
NEWS 9
         Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 10
         Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
         Oct 24 BEILSTEIN adds new search fields
NEWS 11
        Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 12
NEWS 13
         Nov 18
                DKILIT has been renamed APOLLIT
NEWS 14
         Nov 25 More calculated properties added to REGISTRY
NEWS 15
        Dec 04 CSA files on STN
NEWS 16
        Dec 17
                PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 17
         Dec 17 'TOXCENTER enhanced with additional content
NEWS 18
         Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 19
         Jan 29
                 Simultaneous left and right truncation added to COMPENDEX,
                 ENERGY, INSPEC
         Feb 13 CANCERLIT is no longer being updated
NEWS 20
NEWS 21 Feb 24 METADEX enhancements
NEWS 22 Feb 24 PCTGEN now available on STN
NEWS 23 Feb 24 TEMA now available on STN
NEWS 24 Feb 26 NTIS now allows simultaneous left and right truncation
NEWS 25 Feb 26 PCTFULL now contains images
NEWS 26 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 27 Mar 20 EVENTLINE will be removed from STN
NEWS 28 Mar 24
                PATDPAFULL now available on STN
NEWS 29
        Mar 24 Additional information for trade-named substances without
                 structures available in REGISTRY
NEWS 30
        Apr 11
                Display formats in DGENE enhanced
NEWS 31
        Apr 14
                MEDLINE Reload
NEWS 32
        Apr 17
                Polymer searching in REGISTRY enhanced
NEWS 33
        Apr 21
                Indexing from 1947 to 1956 being added to records in CA/CAPLUS
NEWS 34
        Apr 21
                New current-awareness alert (SDI) frequency in
                WPIDS/WPINDEX/WPIX
        Apr 28
NEWS 35
                RDISCLOSURE now available on STN
NEWS 36
        May 05
                Pharmacokinetic information and systematic chemical names
                 added to PHAR
NEWS 37
                MEDLINE file segment of TOXCENTER reloaded
        May 15
NEWS 38
                Supporter information for ENCOMPPAT and ENCOMPLIT updated
        May 15
NEWS 39
        May 16
                CHEMREACT will be removed from STN
NEWS 40
        May 19
                Simultaneous left and right truncation added to WSCA
NEWS 41
        May 19
                RAPRA enhanced with new search field, simultaneous left and
                right truncation
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NEWS EXPRESS April 4 CURRENT WINDOWS VERSION IS V6.01a, CURRENT

MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP), AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003 STN Operating Hours Plus Help Desk Availability General Internet Information Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN NEWS WWW CAS World Wide Web Site (general information)

. Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 15:49:45 ON 02 JUN 2003

=> e reg

NEWS HOURS

NEWS INTER

NEWS LOGIN

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE The EXPAND command is used to look at the index in a file which has an index. This file does not have an index.

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 15:49:57 ON 02 JUN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 1 JUN 2003 HIGHEST RN 523977-56-2 DICTIONARY FILE UPDATES: 1 JUN 2003 HIGHEST RN 523977-56-2

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> e fumitremorgin

| E1 | 4 | FUMITOXIN/BI |
|----|-----|------------------|
| E2 | 1 | FUMITREMORGEN/BI |
| E3 | 36> | FUMITREMORGIN/BI |
| E4 | 3 | FUMJUDAINE/BI |
| E5 | 1 | FUMMITE/BI |
| E6 | 2 | FUMOFICIN/BI |
| E7 | 1 | FUMOFICINAL/BT |

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E8
             1
                   FUMOFICINALINE/BI
E9
             1
                   FUMOFICINAMINE/BI
E10
           145
                   FUMONISIN/BI
             7
E11
                   FUMOSA/BI
                   FUMOSIAVELLANEA/BI
E12
             1
=> s e3
L1
            36 FUMITREMORGIN/BI
=> d 11 30-36
L1
     ANSWER 30 OF 36 REGISTRY COPYRIGHT 2003 ACS
     55387-47-8 REGISTRY
RN
     Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
CN
     1'-(acetyloxy)-4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-
     quinazolinyl]-1',3,4,9'a-tetrahydro-2',2'-dimethyl-, [9'S-
     [9'.alpha.[4S*(R*)],9'a.beta.]]- (9CI) (CA INDEX NAME)
OTHER NAMES:
     Fumitremorgin C acetate
CN
CN
     Tryptoquivaline A acetate
CN
     Tryptoquivaline acetate
MF
     C31 H32 N4 O8
LC
     STN Files:
                  BEILSTEIN*, CA, CAPLUS
         (*File contains numerically searchable property data)
```

```
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
```

L1 ANSWER 31 OF 36 REGISTRY COPYRIGHT 2003 ACS
RN 55387-45-6 REGISTRY
CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
 4-[2-[(1S)-1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl] 1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-, (2S,4R,9'aS)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Spiro[furan-2(5H),9'-[9H]imidazo[1,2-a]indole]-3',5(2'H)-dione,
 4-[2-[1-(acetyloxy)-2-methylpropyl]-4-oxo-3(4H)-quinazolinyl]-1',3,4,9'a-tetrahydro-1'-hydroxy-2',2'-dimethyl-, [9'S-[9'.alpha.[4S*(R*)],9'a.beta.]
]-

2 REFERENCES IN FILE CA (1957 TO DATE)
2 REFERENCES IN FILE CAPLUS (1957 TO DATE)

OTHER NAMES:

CN Fumitremorgin C

CN Tryptoquivaline

CN Tryptoquivaline A
CN Tryptoquivaline C
FS STEREOSEARCH
MF C29 H30 N4 O7
LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA,
CANCERLIT, CAPLUS, EMBASE, MEDLINE, NAPRALERT, TOXCENTER
(*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

12 REFERENCES IN FILE CA (1957 TO DATE)
12 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 32 OF 36 REGISTRY COPYRIGHT 2003 ACS

RN 54009-33-5 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methylbutyl)-12-(2-methylpropyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Tetrahydrofumitremorgin B

CN Tetrahydrolanosulin

FS STEREOSEARCH

MF C27 H37 N3 O5

LC STN Files: BEILSTEIN*, CA, CAPLUS, TOXCENTER (*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1957 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 33 OF 36 REGISTRY COPYRIGHT 2003 ACS
- RN 54009-32-4 REGISTRY
- CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methylbutyl)-12-(2-methyl-1-propenyl)-, [5aR-(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]- (9CI) (CA INDEX NAME)

OTHER NAMES:

- CN 22,23-Dihydrofumitremorgin B
- CN Dihydrofumitremorgin B
- CN Dihydrolanosulin
- FS STEREOSEARCH
- MF C27 H35 N3 O5
- LC STN Files: BEILSTEIN*, CA, CAPLUS, TOXCENTER

(*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1957 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1957 TO DATE)
- L1 ANSWER 34 OF 36 REGISTRY COPYRIGHT 2003 ACS
- RN 12626-18-5 REGISTRY
- CN 5H,12H-3,4-Dioxa-5a,11a,15a-triazacyclooct[lm]indeno[5,6-b]fluorene-11,15(2H,13H)-dione, 1,10,10a,14,14a,15b-hexahydro-10a-hydroxy-7-methoxy-2,2-dimethyl-10-[(3-methyl-2-butenyl)oxy]-5-(2-methyl-1-propenyl)-, (5R,10s,10aR,14aS,15bS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

- CN 5H,12H-3,4-Dioxa-5a,11a,15a-triazacyclooct[lm]indeno[5,6-b]fluorene-11,15(2H,13H)-dione, 1,10,10a,14,14a,15b-hexahydro-10a-hydroxy-7-methoxy-2,2-dimethyl-10-[(3-methyl-2-butenyl)oxy]-5-(2-methyl-1-propenyl)-, [5R-(5.alpha.,10.alpha.,10a.alpha.,14a.alpha.,15b.alpha.)]-
- OTHER NAMES:
- CN Fumitremorgen A
- CN Fumitremorgin A
- FS STEREOSEARCH
- MF C32 H41 N3 O7
- LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAPLUS, DDFU, DRUGU, EMBASE, MEDLINE, NAPRALERT, RTECS*, TOXCENTER, USPATEULI
 - (*File contains numerically searchable property data)

Absolute stereochemistry.

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**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
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24 REFERENCES IN FILE CA (1957 TO DATE)
24 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 35 OF 36 REGISTRY COPYRIGHT 2003 ACS

RN 12626-17-4 REGISTRY

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14-dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3-methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, (5aR,6S,12S,14aS)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 5H,14H-Pyrrolo[1'',2'':4',5']pyrazino[1',2':1,6]pyrido[3,4-b]indole-5,14dione, 1,2,3,5a,6,11,12,14a-octahydro-5a,6-dihydroxy-9-methoxy-11-(3methyl-2-butenyl)-12-(2-methyl-1-propenyl)-, [5aR(5a.alpha.,6.alpha.,12.beta.,14a.alpha.)]-

OTHER NAMES:

CN Fumitremorgin B

CN Lanosulin

CN NA 209B

FS STEREOSEARCH

DR 40451-43-2

MF C27 H33 N3 O5

CI COM

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAPLUS, CASREACT, DDFU, DRUGU, EMBASE, MEDLINE, NAPRALERT, RTECS*, SPECINFO, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

42 REFERENCES IN FILE CA (1957 TO DATE)

1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

42 REFERENCES IN FILE CAPLUS (1957 TO DATE)

L1 ANSWER 36 OF 36 REGISTRY COPYRIGHT 2003 ACS

RN 11100-25-7 REGISTRY

CN Fumitremorgin (9CI) (CA INDEX NAME)

MF Unspecified

CI MAN

LC STN Files: AGRICOLA, BIOBUSINESS, BIOSIS, CA, CAPLUS, TOXCENTER

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

9 REFERENCES IN FILE CA (1957 TO DATE)

2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

9 REFERENCES IN FILE CAPLUS (1957 TO DATE)

=>

=> file caplus
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 19.98 20.19

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 15:56:12 ON 02 JUN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 2 Jun 2003 VOL 138 ISS 23 FILE LAST UPDATED: 1 Jun 2003 (20030601/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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=> e neoplasm
                    NEOPLASIQUES/BI
E2
             8
                   NEOPLASIS/BI
E3
        294242 --> NEOPLASM/BI
E4
           183
                   NÉOPLASMA/BI
E5
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                   NEOPLASMAGEL/BI
Еб
            31
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E7
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The search profile that was entered contains terms or
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=> e cancer
E1 ·
            13
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E3
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E6
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L3
=> s l1
L4
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=> d 14 and 12
L2 IS NOT VALID HERE
For an explanation, enter "HELP DISPLAY".
=> s 14 and112
MISSING OPERATOR L4 ANDLL2
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The search profile that was entered contains terms or

nested terms that are not separated by a logical operator. => s 14 and 12 8 L4 AND L2 => d 15 1-8 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2003 ACS 2002:290794 CAPLUS ΑN DN 136:304053 TI Reversal of multidrug resistance in human colon carcinoma cells using fumitremorgins and diketopiperazines IN Rabindran, Sridhar Krishna; He, Haiyin; Greenberger, Lee Martin PA American Cyanamid Company, USA SO U.S., 19 pp. CODEN: USXXAM DT Patent LΑ English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. PI US 6372775 В1 20020416 US 1999-321182 19990527 US 2002156015 A1 20021024 US 2002-86169 20020228 US 2002169111 US 2002-86132 Α1 20021114 20020228 US 6537964 В1 20030325 US 2002-86170 20020228 US 2003083230 Α1 20030501 US 2002-86133 20020228 PRAI US 1998-109801P Ρ 19980527 US 1999-321182 А3 19990527 OS . MARPAT 136:304053 RE.CNT 72 THERE ARE 72 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT L5 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2003 ACS 2000:824069 CAPLUS AN DN 134:9341 TI A method of improving bioavailability of orally administered drugs, screening for enhancers of such bioavailability and novel pharmaceutical compositions for oral delivery of drugs IN Schellens, Johannes Henricus Matthias; Schinkel, Alfred Hermanus Het Nederlands Kankerinstituut, Neth. PA PCT Int. Appl., 25 pp. SO CODEN: PIXXD2 DTPatent LА English FAN.CNT 2 PATENT NO. KIND DATE APPLICATION NO. DATE ____ _____ -----WO 2000069390 PΙ Α2 20001123 WO 2000-NL331 20000517 WO 2000069390 Α3 20011213 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, W: CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 2000049552 A5 20010205 AU 2000-49552 20000517 EP 1189637 Α2 20020327 EP 2000-931720 20000517 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

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     NL 1999-1012481
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AN
     2000:72225
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DN
     132:216690
TI
     Fumitremorgin C reverses multidrug resistance in cells transfected with
     the breast cancer resistance protein
     Rabindran, Sridhar K.; Ross, Douglas D.; Doyle, L. Austin; Yang, Weidong;
ΑU
     Greenberger, Lee M.
CS
     Oncology and Immunoinflammatory Research, Wyeth-Ayerst Research, Pearl
     River, NY, 10965, USA
SO
     Cancer Research (2000), 60(1), 47-50
     CODEN: CNREA8; ISSN: 0008-5472
     AACR Subscription Office
PB
     Journal
DT
     English
              THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD
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     ANSWER 4 OF 8 CAPLUS COPYRIGHT 2003 ACS
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ΑN
     2000:27209 CAPLUS
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     132:180759
ΤI
     Fumitremorgin C analogs that reverse mitoxantrone resistance in human
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    He, Haiyin; Rabindran, Sridhar G.; Greenberger, Lee M.; Carter, Guy T.
ΑU
     Natural Products Chemistry, Wyeth-Ayerst Research, Pearl River, NY, 10965,
CS
SO
     Medicinal Chemistry Research (1999), 9(6), 424-437
     CODEN: MCREEB; ISSN: 1054-2523
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ΤI
    A multidrug resistance protein associated with antitumor drug resistance
     in breast cancer and a cDNA encoding it
IN
     Ross, Douglas D.; Doyle, L. Austin; Abruzzo, Lynne
PA
    University of Maryland, Baltimore, USA
SO
     PCT Int. Appl., 79 pp.
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                                           APPLICATION NO.
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             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
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RE.CNT 3
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L5
     ANSWER 6 OF 8 CAPLUS COPYRIGHT 2003 ACS
AN
     1999:2922 CAPLUS
     130:177236
DN.
     Reversal of a novel multidrug resistance mechanism in human colon
TI
     carcinoma cells by fumitremorgin C
     Rabindran, Sridhar K.; He, Haiyin; Singh, Maya; Brown, Eileen; Collins,
ΑU
     Karen I.; Annable, Tami; Greenberger, Lee M.
     Oncology and Immunology Research, Wyeth-Ayerst Research, Pearl River, NY,
CS
     10965, USA
     Cancer Research (1998), 58(24), 5850-5858
SO
     CODEN: CNREA8; ISSN: 0008-5472
PB
     AACR Subscription Office
DT
     Journal
LΑ
     English
              THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 51
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L5
     ANSWER 7 OF 8 CAPLUS COPYRIGHT 2003 ACS
AN
     1996:703657 CAPLUS
DN
     126:16546
     Isolation, structure determination and biological activities of novel
```

- TТ mammalian cell cycle inhibitors, spirotryprostatins A & B, tryprostatins A & B and related new diketopiperazine derivatives produced by a fungus, Aspergillus fumigatus
- ΑU Cui, Cheng-Bin; Kakeya, Hideaki; Osada, Hiroyuki
- CS Institute Physical and Chemical Research, Japan
- SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1996), 38th, 49-54 CODEN: TYKYDS
- PB Nippon Kagakkai
- DTJournal
- LΑ Japanese
- L5 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2003 ACS
- AN1992:210833 CAPLUS
- DN 116:210833
- TI Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish
- UΑ Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro
- CS Osaka Univ. Pharm. Sci., Osaka, Japan
- SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS
- DT Journal
- LA Japanese

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ANSWER 7 OF 8 CAPLUS COPYRIGHT 2003 ACS
AN .
     1996:703657 CAPLUS
DN
     126:16546
ΤI
     Isolation, structure determination and biological activities of novel
     mammalian cell cycle inhibitors, spirotryprostatins A & B, tryprostatins A
     & B and related new diketopiperazine derivatives produced by a fungus,
     Aspergillus fumigatus
ΑU
     Cui, Cheng-Bin; Kakeya, Hideaki; Osada, Hiroyuki
CS
     Institute Physical and Chemical Research, Japan
SO
     Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1996), 38th, 49-54
     CODEN: TYKYDS
PΒ
     Nippon Kagakkai
DT
     Journal
LΑ
     Japanese
CC
     10-1 (Microbial, Algal, and Fungal Biochemistry)
     Section cross-reference(s): 1
GI
* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *
AΒ
     A discussion is given on the prodn., isolation, structure detn., and biol.
     activities of the novel Aspergillus fumigatus diketopiperazine alkaloids
     spirotryprostatins A (I) and B, tryprostatins A (II) and B, and
     cyclotryprostatins B and D, the 3 new natural diketopiperazines
     cyclotryprostatins A (III) and C, and demethoxyfumitremorgin C (IV), and
     on several known diketopiperazines, such as fumitremorgin C. Structures
     of the new diketopiperazines were detd. mainly by spectroscopic methods.
    All of the compds. are inhibitors of the G2/M transition of tsFT210 cells
     at micromolar dosages. Some structure-activity relations were obsd.
ST
     Aspergillus diketopiperazine alkaloid cell cycle inhibitor;
     spirotryprostatin mammal cell cycle inhibitor Aspergillus; antitumor
     diketopiperazine alkaloid Aspergillus; tryprostatin mammal cell cycle
     inhibitor Aspergillus; neoplasm inhibitor diketopiperazine deriv
     Aspergillus
ΙT
    Mitosis
        (-G2 transition; diketopiperazine derivs. produced and isolated from
        Aspergillus fumigatus are novel mammalian cell cycle inhibitors)
ΙT
     Interphase (cell cycle)
        (G2-phase, -M transition; diketopiperazine derivs. produced and
        isolated from Aspergillus fumigatus are novel mammalian cell cycle
        inhibitors)
IT
     Structure-activity relationship
      (antimitotic; of diketopiperazines)
     Structure-activity relationship
IT
        (antitumor; of diketopiperazines)
IT
    New natural products
        (cyclotryprostatin A (diketopiperazine))
IT
    New natural products
        (cyclotryprostatin B (diketopiperazine))
ΙT
    New natural products
        (cyclotryprostatin C (diketopiperazine))
IT
    New natural products
        (cyclotryprostatin D (diketopiperazine))
IT
    Antitumor agents
        (diketopiperazine derivs. from Aspergillus fumigatus as)
TΤ
    Aspergillus fumigatus
     Fermentation
        (diketopiperazine derivs. produced and isolated from Aspergillus
```

fumigatus are novel mammalian cell cycle inhibitors) ΙT Molecular structure, natural product (of cyclotryprostatin A (diketopiperazine)) IT Molecular structure, natural product (of cyclotryprostatin B (diketopiperazine)) IT Molecular structure, natural product (of cyclotryprostatin C (diketopiperazine)) ΙT Molecular structure, natural product (of cyclotryprostatin D (diketopiperazine)) IT Alkaloids, biological studies RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (piperazine, dioxo; diketopiperazine derivs. produced and isolated from Aspergillus fumigatus are novel mammalian cell cycle inhibitors) 111427-97-5P, Cyclotryprostatin C IT 111468-06-5P, Cyclotryprostatin A 111768-16-2P 118974-02-0P, Fumitremorgin C. 171864-80-5P, Tryprostatin A 179936-52-8P, Tryprostatin B 182234-25-9P, Spirotryprostatin A 182234-26-0P, Spirotryprostatin B 184305-67-7P, Cyclotryprostatin B 184305-68-8P, Cyclotryprostatin D RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (diketopiperazine derivs. produced and isolated from Aspergillus fumigatus are novel mammalian cell cycle inhibitors) L5ANSWER 8 OF 8 CAPLUS COPYRIGHT 2003 ACS ΑN 1992:210833 CAPLUS DN 116:210833 TI Structures of cytotoxic substances and new quinazoline derivatives produced by a fungus from a saltwater fish ΑU Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika; Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi, Hirofumi; Shingu, Tetsuro Osaka Univ. Pharm. Sci., Osaka, Japan CS SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30 CODEN: TYKYDS DTJournal LA Japanese CC 10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1 GΙ

II

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Fifteen metabolites were isolated from the mycelium and culture filtrate
     of a strain of Aspergillus fumigatus which existed in the gastrointestinal
     tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2,
     fumitermorgin C and gliotoxin exhibited significant cytotoxicity against
     the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C
     COSY and other spectral data for the 5 new metabolites [fumiquinazoline
     (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal
     or moderate cytotoxicity, allowed assignment of their structures contg.
     quinazolone and indoline moieties. The ab. stereostructure of III was
     detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of
     L-(+)-alanine by acid hydrolysis. The stereochem. of the other
     metabolites was established by deriving I and V from IV and other chem.
     behavior.
ST
     Aspergillus fumiquinazoline cytotoxicity structure
TΤ
     Nomenclature, new natural products
        (fumiquinazoline A (quinazoline), from Aspergillus fumigatus)
     Aspergillus fumigatus
TΤ
        (fumiquinazolines from, structure and cytotoxicity of)
IT
     Molecular structure, natural product
        (of fumiquinazoline A (quinazoline), from Aspergillus fumigatus)
IT
     Molecular structure, natural product
        (of fumiquinazoline B (quinazoline), from Aspergillus fumigatus)
IT
     Molecular structure, natural product
        (of fumiquinazoline C (quinazoline), from Aspergillus fumigatus)
IT
     Molecular structure, natural product
        (of fumiquinazoline D (quinazoline), from Aspergillus fumigatus)
ΙT
     Molecular structure, natural product
        (of fumiquinazoline E (quinazoline), from Aspergillus fumigatus)
IT
     Neoplasm inhibitors
        (leukemia, fumiquinazolines as, from Aspergillus fumigatus)
IT
     140715-88-4P
                    140715-89-5P
                                  140715-90-8P
                                                  140852-72-8P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. of)
IT
     67-99-2, Gliotoxin
                          253-82-7D, Quinazoline, derivs.
                                                            12771-72-1,
                    51177-07-2
     Verruculogen
                                 62867-47-4, Fumigaclavine C
     111427-99-7, TR 3
                         111468-06-5
                                       115589-18-9 118974-02-0
       Fumitremorgin C
                         137494-04-3
                                       140715-85-1, Fumiquinazoline A
     140715-86-2, Fumiquinazoline D
                                      140715-87-3, Fumiquinazoline E
     140852-71-7, Fumiquinazoline B
                                      140924-01-2, Fumiquinazoline C
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (structure and cytotoxic activity of, from Aspergillus fumigatus)
=> s 14 and 13
MISSING OPERATOR L4 ANDL3
The search profile that was entered contains terms or
nested terms that are not separated by a logical operator.
=> s 14 and 13
            12 L4 AND L3
=> d 16 1-12
L6
     ANSWER 1 OF 12 CAPLUS COPYRIGHT 2003 ACS
AN
     2003:335139 CAPLUS
DN
     138:332878
TI
     Application of a human multidrug transporter (abcg2) variant as selectable
     marker in gene transfer to progenitor cells and in gene therapy
IN
     Nemet, Katalin; Varadi, Gyorgy; Cervenak, Judit; Ujhelly, Olga; Sarkadi,
     Balazs; Varadi, Andras; Oezvegy, Csilla
PΑ
     Solvo Biotechnology Inc., Hung.
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SO
     PCT Int. Appl., 44 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 2
     PATENT NO.
                      KIND
                             DATE
                                            APPLICATION NO.
                                                              DATE
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                                            -----
     WO 2003035685
ΡI
                       Α1
                             20030501
                                            WO 2002-HU108
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             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
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             RU, TJ, TM
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             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
     WO 2002071073
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                       A3
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         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
             CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI HU 2001-4446
                             20011024
                     Α
     WO 2002-HU15
                       Α
                             20020304
     HU 2002-3435
                       Α
                             20021011
     HU 2001-947
                       Α
                             20010302
RE.CNT 9
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L6
     ANSWER 2 OF 12 CAPLUS COPYRIGHT 2003 ACS
ΑN
     2002:696659 CAPLUS
     137:222100
DN
ΤI
     Improving bioavailability of orally administered drugs, screening for
     enhancers of such bioavailability and oral drug delivery compositions
IN
     Schellens, Johannes Henricus Matthias; Schinkel, Alfred Hermanus
PA
     Netherlands Cancer Institute, Neth.
     U.S. Pat. Appl. Publ., 11 pp., Cont.-in-part of Appl. No. PCT/NL00/00331.
SO
     CODEN: USXXCO
DT
     Patent
LΑ
     English
FAN.CNT 2
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                              DATE
                             -----
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PΙ
    US 2002128282
                       A1
                             20020912
                                            US 2001-988285
                                                              20011119
    WO 2000069390
                       A2
                             20001123
                                            WO 2000-NL331
                                                              20000517
    WO 2000069390
                       A3
                             20011213
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
             ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
             SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA,
             ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
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DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
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PRAI NL 1999-1012066
                             19990517
                        Α
     NL 1999-1012481
                        Α
                             19990630
     WO 2000-NL331
                             20000517
                        A2
L6
     ANSWER 3 OF 12 CAPLUS COPYRIGHT 2003 ACS
ΑN
     2002:696257 CAPLUS
DN
     137:226574
ΤI
     Screening system based on expression of ABCG2 half transporter protein
IN
     Oezvegy, Csilla; Szakacs, Gergely; Varadi, Andras; Nagy, Zoltan
PA
     Solvo Biotechnology Inc., Hung.
SO
     PCT Int. Appl., 26 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 2
     PATENT NO.
                       KIND
                             DATE
                                             APPLICATION NO.
                                                               DATE
PΙ
     WO 2002071073
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                                             WO 2002-HU15
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     WO 2002071073
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                             20030403
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             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
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             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     WO 2003035685
                       A1
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                                             WO 2002-HU108
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             NE, SN, TD, TG
PRAI HU 2001-947
                             20010302
                       Α
     HU 2001-4446
                             20011024
                       Α
     WO 2002-HU15
                        Α
                             20020304
     HU 2002-3435
                        Α
                             20021011
L6
     ANSWER 4 OF 12 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:524344
                  CAPLUS
DN
     138:201203
     Flow cytometric analysis of breast cancer resistance protein
TI
     expression and function
ΑU
     Minderman, Hans; Suvannasankha, Attaya; O'Loughlin, Kieran L.; Scheffer,
     George L.; Scheper, Rik J.; Robey, Robert W.; Baer, Maria R.
CS
     Leukemia Section, Department of Medicine, Roswell Park Cancer Institute,
     Buffalo, NY, USA
SO
     Cytometry (2002), 48(2), 59-65
     CODEN: CYTODQ; ISSN: 0196-4763
PB
     Wiley-Liss, Inc.
DT
     Journal
LΑ
     English
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THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 27 ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L6 ANSWER 5 OF 12 CAPLUS COPYRIGHT 2003 ACS
- ΑN 2002:290794 CAPLUS
- DN 136:304053
- Reversal of multidrug resistance in human colon carcinoma cells using ΤI fumitremorgins and diketopiperazines
- Rabindran, Sridhar Krishna; He, Haiyin; Greenberger, Lee Martin IN
- PA American Cyanamid Company, USA
- SO U.S., 19 pp. CODEN: USXXAM
- DТ
- Patent
- LΑ English
- FAN. CNT 1

| L'UIA • | TAN CIVI I | | | | | | |
|----------|----------------------------|------------|----------|-----------------|----------|--|--|
| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE | | |
| | | | | | | | |
| ΡI | US 6372775 | В1 | 20020416 | US 1999-321182 | 19990527 | | |
| | US 2002156015 | A1 | 20021024 | US 2002-86169 | 20020228 | | |
| | us 2002169111 [.] | A1 | 20021114 | US 2002-86132 | 20020228 | | |
| | US 6537964 | В1 | 20030325 | US 2002-86170 | 20020228 | | |
| | US 2003083230 | A1 | 20030501 | US 2002-86133 | 20020228 | | |
| PRAI | US 1998-109801P | P | 19980527 | | | | |
| | US 1999-321182 | A 3 | 19990527 | | | | |
| \cap S | MARRAT 136.30/05 | 2 | | | | | |

- MARPAT 136:304053
- THERE ARE 72 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 72 ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L6 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2003 ACS
- 2002:46901 CAPLUS AN
- 137:125308 DN
- TΙ Solid phase synthesis of fumitremorgin-type and other indole alkaloids based on cyclization/cleavage strategy
- ΑU van Loevezijn, Arnold; Rodenko, Boris; Sorm, Willem P.; van Maarseveen, Jan H.; Stegman, Karel; Visser, Geb M.; van Delft, Floris L.; Koomen, Gerrit-Jan
- Laboratory of Organic Chemistry, Institute for Molecular Chemistry, CS University of Amsterdam, Amsterdam, NL-1018 WS, Neth.
- SO Innovation and Perspectives in Solid Phase Synthesis & Combinatorial Libraries: Peptides, Proteins and Nucleic Acids--Small Molecule Organic Chemistry Diversity, Collected Papers, International Symposium, 6th, York, United Kingdom, Aug. 31-Sept. 4, 1999 (2001), Meeting Date 1999, 367-370. Editor(s): Epton, Roger. Publisher: Mayflower Scientific Ltd., Kingswinford, UK. CODEN: 69CEGV; ISBN: 0-9515735-3-5
- DT Conference
- LA English
- RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- ANSWER 7 OF 12 CAPLUS COPYRIGHT 2003 ACS L6
- 2001:487684 CAPLUS ΑN
- DN 135:328480
- TI Functional Characterization of the Human Multidrug Transporter, ABCG2, Expressed in Insect Cells
- ΑU Ozvegy, Csilla; Litman, Thomas; Szakacs, Gergely; Nagy, Zoltan; Bates, Susan; Varadi, Andras; Sarkadi, Balazs
- CS Institute of Enzymology, Biological Research Center, Hungarian Academy of Sciences, Budapest, H-1113, Hung.
- SO Biochemical and Biophysical Research Communications (2001), 285(1),
 - CODEN: BBRCA9; ISSN: 0006-291X

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PB
     Academic Press
     Journal
LΑ
     English
RE.CNT 32
               THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L6
     ANSWER 8 OF 12 CAPLUS COPYRIGHT 2003 ACS
ΑN
     2000:872654 CAPLUS
DN
     134:216800
ΤI
     Inhibition of BCRP-mediated drug efflux by fumitremorgin-type indolyl
     diketopiperazines
ΑU
     van Loevezijn, A.; Allen, J. D.; Schinkel, A. H.; Koomen, G.-J.
CS
     Institute of Molecular Chemistry, Laboratory of Organic Chemistry,
     University of Amsterdam, Amsterdam, NL-1018 WS, Neth.
     Bioorganic & Medicinal Chemistry Letters (2000), Volume Date 2001, 11(1),
SO
     29-32
     CODEN: BMCLE8; ISSN: 0960-894X
     Elsevier Science Ltd.
PB
DT
     Journal
LA
     English
RE.CNT 23
               THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 9 OF 12 CAPLUS COPYRIGHT 2003 ACS
Ь6
AN
     2000:824069 CAPLUS
DN
     134:9341
     A method of improving bioavailability of orally administered drugs,
TI
     screening for enhancers of such bioavailability and novel pharmaceutical
     compositions for oral delivery of drugs
IN
     Schellens, Johannes Henricus Matthias; Schinkel, Alfred Hermanus
PA
     Het Nederlands Kankerinstituut, Neth.
     PCT Int. Appl., 25 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 2
     PATENT NO.
                       KIND DATE
                                            APPLICATION NO. DATE
     ______
                       ____
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PΙ
     WO 2000069390
                       A2
                             20001123
                                             WO 2000-NL331
                                                               20000517
     WO 2000069390
                       A3
                             20011213
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
              ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
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                             19990630
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L6
     2000:72225 CAPLUS
AN
DN
     132:216690
ΤI
     Fumitremorgin C reverses multidrug resistance in cells transfected with
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the breast cancer resistance protein
ΑU
     Rabindran, Sridhar K.; Ross, Douglas D.; Doyle, L. Austin; Yang, Weidong;
     Greenberger, Lee M.
CS
     Oncology and Immunoinflammatory Research, Wyeth-Ayerst Research, Pearl
     River, NY, 10965, USA
SO
     Cancer Research (2000), 60(1),
     CODEN: CNREA8; ISSN: 0008-5472
PB
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DT
     Journal
LΑ
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              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L6
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AN
     1999:511178 CAPLUS
DN
     131:143080
     A multidrug resistance protein associated with antitumor drug resistance
TI
     in breast cancer and a cDNA encoding it
     Ross, Douglas D.; Doyle, L. Austin; Abruzzo, Lynne
IN
     University of Maryland, Baltimore, USA
SO
     PCT Int. Appl., 79 pp.
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L6
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AN
     1999:173577 CAPLUS
DN
     131:13477
TI
     Multiple mechanisms confer drug resistance to mitoxantrone in the human
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AU
     Hazlehurst, Lori A.; Foley, Nils E.; Gleason-Guzman, Mary C.; Hacker,
     Miles P.; Cress, Anne E.; Greenberger, Lee W.; De Jong, Mariska C.;
     Dalton, William S.
CS
     Department of Biochemistry, Pharmacology, and Internal Medicine, H. Lee
     Moffitt Cancer Center, University of South Florida, Tampa, FL, 33612, USA
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SO

Cancer Research (1999), 59(5), 1021-1028

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CODEN: CNREA8; ISSN: 0008-5472
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LA
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=> d 18 1-3
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AN
     2002:290794 CAPLUS
DN
     136:304053
TI
     Reversal of multidrug resistance in human colon carcinoma cells
     using fumitremorgins and diketopiperazines
IN
     Rabindran, Sridhar Krishna; He, Haiyin; Greenberger, Lee Martin
PA
     American Cyanamid Company, USA
SO
     U.S., 19 pp.
     CODEN: USXXAM
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L8
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     2000:27209 CAPLUS
ΑN
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     132:180759
TI
     Fumitremorgin C analogs that reverse mitoxantrone resistance in human
     colon carcinoma cells
     He, Haiyin; Rabindran, Sridhar G.; Greenberger, Lee M.; Carter, Guy T.
ΑU
CS
     Natural Products Chemistry, Wyeth-Ayerst Research, Pearl River, NY, 10965,
```

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SO
     Medicinal Chemistry Research (1999), 9(6), 424-437
     CODEN: MCREEB; ISSN: 1054-2523
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AN
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DN
     130:177236
TI 
     Reversal of a novel multidrug resistance mechanism in human colon
     carcinoma cells by fumitremorgin C
ΑU
     Rabindran, Sridhar K.; He, Haiyin; Singh, Maya; Brown, Eileen; Collins,
     Karen I.; Annable, Tami; Greenberger, Lee M.
     Oncology and Immunology Research, Wyeth-Ayerst Research, Pearl River, NY,
CS
     10965, USA
SO
     Cancer Research (1998), 58(24), 5850-5858
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LΑ
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RE.CNT
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     131:143080
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    A multidrug resistance protein associated with antitumor drug resistance
     in breast cancer and a cDNA encoding it
IN
    Ross, Douglas D.; Doyle, L. Austin; Abruzzo, Lynne
PA
    University of Maryland, Baltimore, USA
SO
    PCT Int. Appl., 79 pp.
    CODEN: PIXXD2
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APPLICATION NO.



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     Structures of cytotoxic substances and new quinazoline derivatives
ΤI
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ΑU
     Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika;
     Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi,
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CS
     Osaka Univ. Pharm. Sci., Osaka, Japan
SO
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DT
     Journal
LА
     Japanese
=> d 110 2 all
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ΑN
     1992:210833 CAPLUS
DN
     116:210833
TI
     Structures of cytotoxic substances and new quinazoline derivatives
     produced by a fungus from a saltwater fish
     Numata, Atsushi; Takahashi, Chika; Miyamoto, Tamie; Matsushita, Tomochika;
ΑU
     Kawai, Kenzo; Usami, Yoshihide; Matsumura, Eiko; Inoue, Masatoshi; Ohishi,
     Hirofumi; Shingu, Tetsuro
CS
     Osaka Univ. Pharm. Sci., Osaka, Japan
SO
     Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1991), 33rd, 723-30
     CODEN: TYKYDS
DT
     Journal
LA
     Japanese
CC
     10-1 (Microbial, Algal, and Fungal Biochemistry)
     Section cross-reference(s): 1
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ĠΪ

AB Fifteen metabolites were isolated from the mycelium and culture filtrate of a strain of Aspergillus fumigatus which existed in the gastrointestinal tract of the saltwater fish Pseudolabrus japonicus. Among them, TR-2, fumitermorgin C and gliotoxin exhibited significant cytotoxicity against the cultured P-388 lymphocytic leukemia cells. Anal. of long range 1H-13C COSY and other spectral data for the 5 new metabolites [fumiquinazoline (AFQ-A) (I), -B (II)), -C (III), -D (IV) and -E (V)], exhibiting marginal or moderate cytotoxicity, allowed assignment of their structures contg. quinazolone and indoline moieties. The ab. stereostructure of III was detd. on the basis of x-ray crystallog. anal. as well as of the prodn. of L-(+)-alanine by acid hydrolysis. The stereochem. of the other metabolites was established by deriving I and V from IV and other chem. behavior. ST Aspergillus fumiquinazoline cytotoxicity structure IT Nomenclature, new natural products (fumiquinazoline A (quinazoline), from Aspergillus fumigatus) IT Aspergillus fumigatus (fumiquinazolines from, structure and cytotoxicity of) IT Molecular structure, natural product (of fumiquinazoline A (quinazoline), from Aspergillus fumigatus) IT Molecular structure, natural product (of fumiquinazoline B (quinazoline), from Aspergillus fumigatus) IT Molecular structure, natural product (of fumiquinazoline C (quinazoline), from Aspergillus fumigatus) Molecular structure, natural product ΙT (of fumiquinazoline D (quinazoline), from Aspergillus fumigatus) IT Molecular structure, natural product (of fumiquinazoline E (quinazoline), from Aspergillus fumigatus) IT Neoplasm inhibitors (leukemia, fumiquinazolines as, from Aspergillus fumigatus) IT 140715-88-4P 140715-89-5P 140715-90-8P 140852-72-8P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) IT 67-99-2, Gliotoxin 253-82-7D, Quinazoline, derivs. 12771-72-1, 51177-07-2 62867-47-4, Fumigaclavine C Verruculogen 74149-38-5 111427-99-7, TR 3 111468-06-5 115589-18-9 **118974-02-0** , Fumitremorgin C 137494-04-3 140715-85-1, Fumiquinazoline A 140715-86-2, Fumiquinazoline D 140715-87-3, Fumiquinazoline E 140852-71-7, Fumiquinazoline B 140924-01-2, Fumiquinazoline C RL: BAC (Biological activity or effector, except adverse); BSU (Biological

(structure and cytotoxic activity of, from Aspergillus fumigatus)

study, unclassified); BIOL (Biological study)

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L6 12 S L4 AND L3

E CARCINOMA

L7 100333 S E3

L8 3 S L7 AND L4

E LEUKEMIA

L9 77060 S E3

L10 2 S L9 AND L4

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1996:96400 CAPLUS DN 124:193617 ΤI Non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line Naito, Seiji; Hasegawa, Shuji; Yokomizo, Akira; Koga, Hirofumi; Kotoh, AU Shuji; Kuwano, Michihiko; Kumazawa, Joichi CS Fac. Medicine, Kyushu Univ., Fukuoka, 812, Japan SO Japanese Journal of Cancer Research (1995), 86(11), 1112-18 CODEN: JJCREP; ISSN: 0910-5050 PB Japanese Cancer Association DT Journal LΑ English CC 1-6 (Pharmacology) AB A human bladder cancer cell line resistant to adriamycin (ADM), T24/ADM9 has been established in vitro by exposing T24 parent cells to progressively higher concns. of the drug over a period of 12 mo. The T24/ADM9 cells were 9 times more resistant to ADM than the T24 parent, and showed various degrees of cross-resistance to an ADM deriv., vinca alkaloids and a DNA topoisomerase II (Topo II)-targeting agent, etoposide. No significant difference was obsd. in the cellular accumulation of ADM between the T24/ADM9 and T24 parent cells. A Northern blot anal. showed an overexpression of multidrug resistance -assocd. protein (MRP) mRNA, but no overexpression of multidrug resistance-1 (MDR1) mRNA was obsd. in the T24/ADM9 cells. A flow cytometric anal. showed that the MDR1 gene product, P-glycoprotein (Pgp), is not expressed on the T24/ADM9 cells. T24/ADM9 showed approx. the parental level of DNA Topo II catalytic activity. In Western blot and Northern blot analyses, however, the cellular level of DNA Topo II was apparently much lower in T24/ADM9 than in the T24 parent. Thus, these results suggest that a decreased cellular level of DNA Topo II and an overexpression of MRP gene may be responsible for the expression of an MDR phenotype in the T24/ADM9 cells and that such non-Pgp-mediated, atypical MDR may develop in bladder cancer treated with chemotherapy including ADM. ST atypical multidrug resistant bladder cancer cell ΙŤ Proteins, specific or class RL: BSU (Biological study, unclassified); BIOL (Biological study) (multidrug resistance; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Gene, animal RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line in relation to overexpression of MRP gene) IT Neoplasm inhibitors (bladder carcinoma, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) ΙT Drug resistance (multi-, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT (neoplasm, carcinoma, inhibitors, non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line) IT 50-07-7, Mitomycin C 51-21-8, 5 Fluorouracil 57-22-7, Vincristine 865-21-4, Vinblastine **15663-27-1**, Cisplatin 33419-42-0, 56420-45-2, Epirubicin RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (cross-resistance; non-P-glycoprotein-mediated atypical

multidrug resistance in a human bladder cancer cell line)

IT 142805-56-9, DNA topoisomerase II

RL: BOC (Biological occurrence); BSU (Biological study, unclassified);

BIOL (Biological study); OCCU (Occurrence)

(non-P-glycoprotein-mediated atypical multidrug resistance in

a human bladder cancer cell line in relation to decreased level of DNA

Topo II)

IT 25316-40-9, Adriamycin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(resistance to; non-P-glycoprotein-mediated atypical multidrug resistance in a human bladder cancer cell line)

=>

· DN 126:16546 TI Isolation, structure determination and biological activities of novel mammalian cell cycle inhibitors, spirotryprostatins A & B, tryprostatins A & B and related new diketopiperazine derivatives produced by a fungus, Aspergillus fumigatus Cui, Cheng-Bin; Kakeya, Hideaki; Osada, Hiroyuki ΑU Institute Physical and Chemical Research, Japan CS SO Tennen Yuki Kagobutsu Toronkai Koen Yoshishu (1996), 38th, 49-54 CODEN: TYKYDS PΒ Nippon Kagakkai DTJournal LA Japanese .CC 10-1 (Microbial, Algal, and Fungal Biochemistry) Section cross-reference(s): 1 GΙ

. * STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB A discussion is given on the prodn., isolation, structure detn., and biol. activities of the novel Aspergillus fumigatus diketopiperazine alkaloids spirotryprostatins A (I) and B, tryprostatins A (II) and B, and cyclotryprostatins B and D, the 3 new natural diketopiperazines cyclotryprostatins A (III) and C, and demethoxyfumitremorgin C (IV), and on several known diketopiperazines, such as fumitremorgin C. Structures of the new diketopiperazines were detd. mainly by spectroscopic methods. All of the compds. are inhibitors of the G2/M transition of tsFT210 cells at micromolar dosages. Some structure-activity relations were obsd.

ST Aspergillus diketopiperazine alkaloid cell cycle inhibitor; spirotryprostatin mammal cell cycle inhibitor Aspergillus; antitumor diketopiperazine alkaloid Aspergillus; tryprostatin mammal cell cycle inhibitor Aspergillus; neoplasm inhibitor diketopiperazine deriv Aspergillus

IT Mitosis

(-G2 transition; diketopiperazine derivs. produced and isolated from Aspergillus fumigatus are novel mammalian cell cycle inhibitors)

IT Interphase (cell cycle)

(G2-phase, -M transition; diketopiperazine derivs. produced and isolated from Aspergillus fumigatus are novel mammalian cell cycle inhibitors)

IT Structure-activity relationship

(antimitotic; of diketopiperazines)

IT Structure-activity relationship

(antitumor; of diketopiperazines)

IT New natural products

(cyclotryprostatin A (diketopiperazine))

IT New natural products

(cyclotryprostatin B (diketopiperazine))

IT New natural products

(cyclotryprostatin C (diketopiperazine))

IT New natural products

(cyclotryprostatin D (diketopiperazine))

IT Antitumor agents

(diketopiperazine derivs. from Aspergillus fumigatus as)

IT Aspergillus fumigatus

Fermentation

(diketopiperazine derivs. produced and isolated from Aspergillus fumigatus are novel mammalian cell cycle inhibitors)

IT Molecular structure, natural product

(of cyclotryprostatin A (diketopiperazine))

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IT
    Molecular structure, natural product
        (of cyclotryprostatin B (diketopiperazine))
IT
    Molecular structure, natural product
        (of cyclotryprostatin C (diketopiperazine))
IT
    Molecular structure, natural product
        (of cyclotryprostatin D (diketopiperazine))
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     (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
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       Aspergillus fumigatus are novel mammalian cell cycle inhibitors)
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     111427-97-5P, Cyclotryprostatin C
                                         111468-06-5P, Cyclotryprostatin A
     111768-16-2P 118974-02-0P, Fumitremorgin C.
     171864-80-5P, Tryprostatin A
                                    179936-52-8P, Tryprostatin B
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    182234-25-9P, Spirotryprostatin A
     184305-67-7P, Cyclotryprostatin B
                                         184305-68-8P, Cyclotryprostatin D
    RL: BAC (Biological activity or effector, except adverse); BPN
     (Biosynthetic preparation); BSU (Biological study, unclassified); PRP
     (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL
   (Biological study); PREP (Preparation); USES (Uses)
        (diketopiperazine derivs. produced and isolated from Aspergillus
       fumigatus are novel mammalian cell cycle inhibitors)
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